

# **James A. Haley Veterans' Hospital Antimicrobial Guide: 2020/2021**

*Inpatient Edition*



# **ASP**

**Antimicrobial Stewardship  
Program**

**Provided by:**

Acute Care Section, Pharmacy Service

Infectious Diseases Section, Medical Service

Microbiology Section, Pathology and Laboratory Service

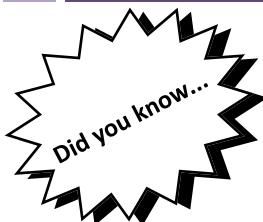
**Approved by:**

Pharmacy and Therapeutics Committee

Antimicrobial Stewardship Subcommittee

**Updated October 2020**

# NOTES PAGE



Updated CLSI Interpretive Categories and MIC Breakpoints - M100 S30

Agent	Organism	S	I	R
Ciprofloxacin	<i>Enterobacteriaceae</i>	$\leq 0.25$	0.5	$\geq 1$
Ciprofloxacin	<i>Pseudomonas aeruginosa</i>	$\leq 0.5$	1	$\geq 2$
Levofloxacin	<i>Enterobacteriaceae</i>	$\leq 0.5$	1	$\geq 2$
Levofloxacin	<i>Pseudomonas aeruginosa</i>	$\leq 1$	2	$\geq 4$

Ciprofloxacin and levofloxacin breakpoints have changed for *Pseudomonas aeruginosa* and *Enterobacteriaceae* (such as *Escherichia*, *Klebsiella*, *Enterobacter*, *Serratia*, *Citrobacter*, *Proteus* spp.) These new breakpoints are substantially lower than previous breakpoints and are NOT yet reflected in CPRS.

For example, an *E. coli* isolate with a ciprofloxacin MIC of 1 will be reported in CPRS as 'S' for susceptible (based on prior breakpoints). However, this is now considered 'R' or resistant and an alternative therapy should be utilized.

In addition, the *Pseudomonas aeruginosa* breakpoints are based on high-dose therapy (ciprofloxacin 400 mg IV Q8H, ciprofloxacin 750 mg PO Q12H, and levofloxacin 750 mg IV/PO Q24H) so be sure to use these higher anti-*Pseudomonal* doses when necessary.

You can read more about minimum inhibitory concentrations (MICs) and interpreting a culture and susceptibility report on page 17. You can also access the full CLSI document here: <http://em100.edaptivedocs.net/Login.aspx>

# TABLE OF CONTENTS

Disease State Overview	
▪ Urinary tract infections algorithm	1-2
▪ Asymptomatic bacteriuria	3
▪ Symptomatic urinary tract infection	4
▪ Community acquired pneumonia	5
▪ Hospital and ventilator associated pneumonia	6
▪ Diabetic foot infections	7
▪ <i>C. difficile</i> infection	8
▪ Infective endocarditis	9-10
▪ Neutropenic prophylaxis	11
▪ Febrile neutropenia	12
▪ Bacterial meningitis	13
▪ Skin and soft tissue infections	14
▪ Influenza	15
▪ Bacteremia	16
Choosing an antibiotic	17
Renal dosing overview	18
Renal dose adjustments – IV antimicrobials	19
Dosing in dialysis	20
Renal dose adjustments – PO antimicrobials	21
Overview of common antiretroviral therapy	22
Alternative dosing (cefepime, meropenem, & piperacillin/tazobactam)	23
Guide to home IV antibiotics	24
Vancomycin dosing/monitoring	25
Aminoglycoside dosing/monitoring – conventional	26
Aminoglycoside dosing/monitoring – extended interval	27-28
Antibiogram (2018 isolates)	29-30
IV to PO conversion	31
Pneumococcal vaccine algorithm	32
Penicillin allergies and $\beta$ -lactam cross reactivity	33-34
Hospital precautions	35
Antibiotic spectrum of activity – at a glance	36
Antibiotic spectrum of activity	37-38
Available antimicrobials at JAHVH	39-40

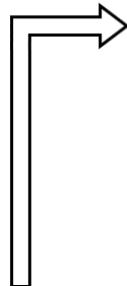
Note, the information provided in this booklet is a summary of published guidelines, literature, and accepted practices as they relate to the JAHVH formulary. It is provided as a reference and should not be used in lieu of clinical decision making. Guidelines may have been updated since the time of printing. For the most complete and up-to-date information, please access the literature directly. All empiric doses recommended herein assume normal renal function.

# URINARY TRACT INFECTIONS ALGORITHM

1-2



Pyuria, cloudy urine, foul smell, or positive urinalysis are NOT symptoms of a urinary tract infection (UTI) and are NOT indications for antibiotic therapy



Does the patient have any urinary signs or symptoms consistent with the following modified diagnostic criteria?	
Cystitis	<p>Urinalysis available:</p> <ul style="list-style-type: none"><li>At least 1 of the following: frequency, urgency, dysuria, or suprapubic pain AND</li><li>At least 1 of the following findings on UA: positive leukocyte esterase or pyuria</li></ul> <p>Urinalysis not available:</p> <ul style="list-style-type: none"><li>At least 2 of the following: frequency, urgency, dysuria, or suprapubic pain</li></ul>
Pyelonephritis	<ul style="list-style-type: none"><li>Fever (<math>&gt;100.4^{\circ}\text{F}</math>) AND</li><li>At least 1 of the following: flank pain or costovertebral angle (CVA) tenderness AND</li><li>No indwelling urinary catheter</li></ul>
Catheter-associated UTI	<ul style="list-style-type: none"><li>Indwelling urinary catheter AND</li><li>At least 1 of the following: fever (<math>&gt;100.4^{\circ}\text{F}</math>), rigors, flank pain, CVA tenderness, acute hematuria, or pelvic discomfort</li></ul>

Follow up results of urinalysis and urine culture. Continue to follow algorithm on next page

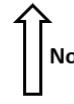
Send urinalysis AND urine culture



Do NOT send urine culture unless patient is pregnant or scheduled to undergo invasive urologic intervention



Does a non-UTI diagnosis likely account for the symptoms?



Work up other cause

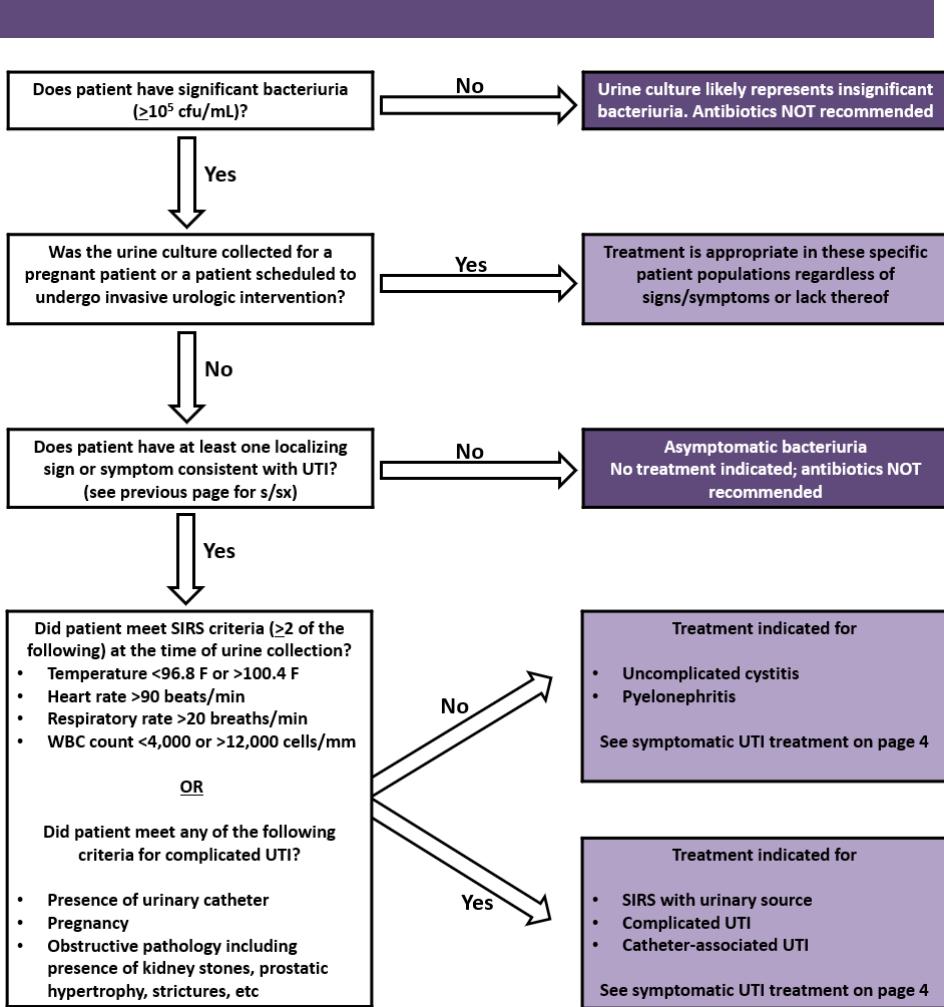


## Urinary Tract Infection (UTI) Diagnosis = Significant bacteriuria ( $\geq 10^5$ ) AND urinary signs/symptoms

- Female:* Two consecutive voided urine specimens with identification of same bacterial strain in quantities  $\geq 100,000 \text{ CFU/mL}$
- Male:* Single, clean-catch voided urine specimen with one bacterial species identified in quantities  $\geq 100,000 \text{ CFU/mL}$
- Catheterized patient:* Single urine specimen with one bacterial species identified in quantities  $\geq 100,000 \text{ CFU/mL}$

## Did You Know?

- Our lab does not reflex urinalysis and urine culture; these tests need to be ordered separately
- Urinalysis results are available quickly while a urine culture may take several days
- UTI cannot be diagnosed based on urinalysis results alone



References for pages 1-4

Algorithm adapted from Antimicrobial Stewardship Task Force UTI Diagnosis and Treatment Algorithm available on VA Antimicrobial Stewardship National Sharepoint site

Nicolle LE et al. Clinical Practice Guideline for the Management of Asymptomatic Bacteriuria: 2019 Update by the Infectious Diseases Society of America. *Clinical Infectious Diseases* 2019; 68(10):83-110

Gupta K et al. International clinical practice guidelines for the treatment of acute uncomplicated cystitis and pyelonephritis in women: A 2010 update by the Infectious Diseases Society of America and the European Society for Microbiology and Infectious Diseases. *Clinical Infectious Diseases* 2010; 52:6e103-20

Hooton TM et al. Diagnosis, Prevention, and Treatment of Catheter-Associated Urinary Tract Infection in Adults:2009 International Clinical Practice Guidelines from the Infectious Diseases Society of America. *Clinical Infectious Diseases* 2010; 50:625-663

"FDA Updates Warnings for Fluoroquinolone Antibiotics" Available at:

<https://www.fda.gov/newsevents/newsroom/pressannouncements/ucm513183.htm> Accessed May 2019

# ASYMPTOMATIC BACTERIURIA (ASB)

3

## Definition

The presence of one or more species of bacteria growing in the urine at a colony count of  $\geq 10^5$  CFU/mL, irrespective of the presence of pyuria, in the absence of signs or symptoms attributable to urinary tract infection (UTI)

- High Prevalence of Asymptomatic Bacteriuria**
- The bladder is frequently colonized with bacteria
  - A positive urinalysis or culture without symptoms should be considered **colonization, NOT infection**
  - Treatment of ASB is **NOT recommended\***

\*See below for select indications requiring treatment for ASB



- Cost of Unnecessary Rx & Missed Diagnosis**
- Drug/drug interactions
  - Renal and other complications
  - Drug allergies
  - C. difficile* infection
  - Missing the real diagnosis
  - No improvement in "UTI"
  - Increased risk of **developing resistant organisms**

Fiction	Fact
Bacteria on urinalysis always indicates UTI	<ul style="list-style-type: none"><li>Urinalyses are often contaminated in the elderly</li></ul>
Urinalysis should be ordered as a screening test	<ul style="list-style-type: none"><li>The high prevalence of asymptomatic bacteriuria means these tests <b>SHOULD NOT</b> be done routinely</li><li>Only screen for ASB in pregnant women and before invasive urologic procedures with risk of mucosal bleeding</li><li>Test only when UTI symptoms are present</li></ul>
An abnormal urinalysis is a good explanation for weakness, fatigue, or mental status changes	<ul style="list-style-type: none"><li>Many elderly patients have asymptomatic bacteriuria</li><li>Seek other causes to ensure appropriate diagnosis; consider using the PINCH-ME approach: <math>P - \text{pain}</math>      <math>M - \text{other medication}</math> <math>I - \text{other infection}</math>      <math>E - \text{environment change}</math> <math>N - \text{poor nutrition}</math> <math>C - \text{constipation}</math> <math>H - \text{poor hydration}</math> • Hydrate if not contraindicated</li></ul>
Pyuria always indicates UTI	<ul style="list-style-type: none"><li>Pyuria is common in patients with ASB – as high as 100% in patients with long-term catheters</li></ul>
Patients with pyuria and bacteriuria need antibiotics	<ul style="list-style-type: none"><li>Neither pyuria nor bacteriuria indicate a need for antibiotics. With rare exceptions treat ONLY symptomatic patients</li></ul>

## Select Populations Who Should NOT be Screened for ASB

- Functionally impaired older adults residing in the community or long-term care facility
- Diabetic patients
- Solid organ transplant patients or renal transplant patients who have had renal transplant surgery  $>1$  month prior
- Patients with neutropenia
- Spinal cord injury patients
- Patients with indwelling catheters
- Patients undergoing surgery for artificial urine sphincter or penile prosthesis implantation

## Select Patients for Which ASB Should be Screened & Treated

- Pregnant women; recommended duration 4-7 days depending on regimen chosen
- Patients undergoing endoscopic urologic procedures associated with mucosal trauma; recommended duration of 1-2 doses initiated 30-60 minutes prior to procedure

# SYMPTOMATIC URINARY TRACT INFECTION

4

## Definitions

- **Cystitis:** lower urinary tract (bladder) infection
- **Pyelonephritis:** upper urinary tract (kidney) infection
- **Catheter-associated UTI (CA-UTI):** urinary tract infection associated with indwelling urethral, indwelling suprapubic, or intermittent catheterization

## Diagnosis

See modified diagnostic criteria on page 1

## Common Causative Organisms

*E. coli*, *Proteus spp.*, *Klebsiella spp.*, *Pseudomonas aeruginosa*

## Empiric Treatment

- When available, empiric treatment should be based off results of prior urine cultures
- Use pathogen-specific antibiotic therapy once urine culture results are available

	Preferred	Alternative
Cystitis (lower UTI)	<ul style="list-style-type: none"><li>• TMP/SMZ 1 DS tablet PO q12h</li><li>• nitrofurantoin* 100 mg PO q12h</li></ul>	<ul style="list-style-type: none"><li>• amoxicillin/clavulanate 875/125 mg PO q12h</li><li>• cefdinir 300 mg PO q12h</li><li>• fosfomycin§ 3 g PO q72h</li></ul>
Pyelonephritis (upper UTI)	<ul style="list-style-type: none"><li>• ceftriaxone 1 g IV q24h</li><li>• cefepime 1 g IV q12h (if mild-moderate)</li><li>• cefepime 1 g IV q6h (if severe)</li></ul>	• ciprofloxacin 400 mg IV q8-12h
CA-UTI	Change catheter then choose one of the above treatment options based on site of infection (lower vs upper urinary tract)	

\*Nitrofurantoin should not be used for treatment of upper urinary tract infections because it concentrates in the urine. Although the nitrofurantoin package insert does not recommend use in patients with CrCl <60 ml/min, there is literature supporting safe nitrofurantoin use for short durations ( $\leq 7$  days) in patients with CrCl > 30 ml/min

§See Clinical Pearls section below

**NOTE:** As of July 2016, the FDA recommends fluoroquinolones be reserved for use in patients who have no other treatment options for uncomplicated UTIs because the risk of serious side effects generally outweighs the benefits. Additionally, local resistance rates of levofloxacin and ciprofloxacin to common causative organisms are high.

**\*\*AVOID fluoroquinolones for the treatment of uncomplicated urinary tract infections\*\***

## Suggested Treatment Duration

Uncomplicated cystitis (males)	7 – 10 days, depending on antibiotic class
Uncomplicated cystitis (females)	3 – 7 days, depending on antibiotic class
Pyelonephritis	10 – 14 days
SIRS with urinary source	10 – 14 days
Complicated urinary tract infection	10 – 14 days
Catheter associated urinary tract infection	7 days if prompt symptom resolution 10 – 14 days if delayed response

## Clinical Pearls

- Oral fosfomycin may be an option for some patients with uncomplicated cystitis caused by either ESBL+ *E. coli* or select Gram negatives with contraindications to other oral options. Fosfomycin should not be used for the treatment of upper urinary tract infections. This medication is nonformulary and requires a Prior Authorization Drug Request (PADR) consult. Please refer to local criteria-for-use (CFU).
- When using fluoroquinolones to treat *Pseudomonas aeruginosa*, anti-Pseudomonal doses should be used:
  - Ciprofloxacin 750 mg PO q12h or ciprofloxacin 400 mg IV q8h
  - Levofloxacin 750 mg PO/IV q24h

# COMMUNITY ACQUIRED PNEUMONIA

5

## Diagnosis

Requires both the presence of clinical features (cough, fever, sputum production, pleuritic chest pain) AND chest infiltrate demonstrated on imaging

## Common Etiologic Agents

*Streptococcus pneumoniae*, *Mycoplasma pneumoniae*, *Haemophilus influenzae*, *Staphylococcus aureus*, *Chlamydophila pneumoniae*, *Moraxella catarrhalis*, and respiratory viruses

## Clinical Pearls

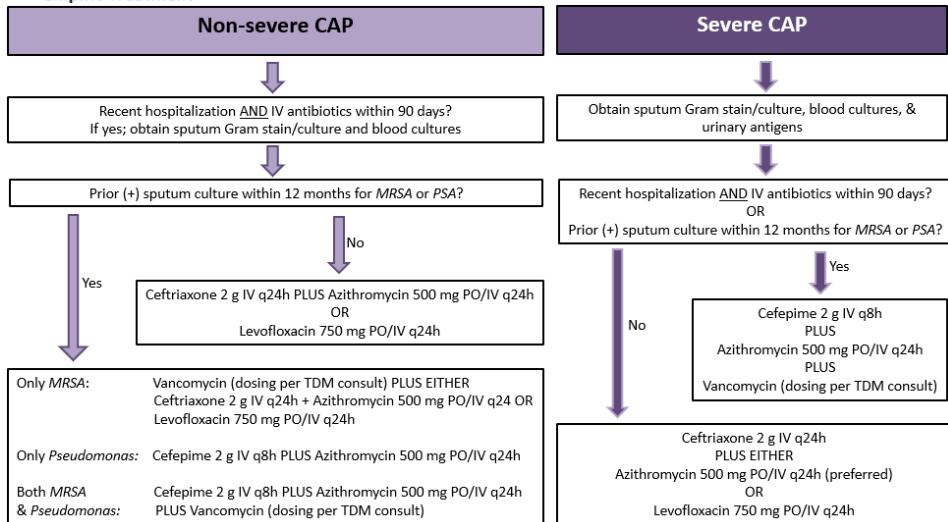
- Obtain sputum cultures before starting antimicrobial therapy
- Consider PSI scores to help identify appropriate treatment strategies; outpatient vs. inpatient
- Signs and symptoms of CAP may be lacking or altered in elderly patients
- Cough and chest X-ray abnormalities may take up to 6 weeks to improve and are NOT a valid reason for extended antibiotic courses

## Classification

**Non-severe CAP**      Patient meets no major and 2 or fewer minor criteria below  
**Severe CAP**      Patient meets at least 1 major or 3 minor criteria below

Major Criteria	Minor Criteria	
Septic shock with need for vasopressors	RR $\geq 30$ breaths/min	PaO <sub>2</sub> /FiO <sub>2</sub> ratio $\leq 250$
Respiratory failure requiring mechanical ventilation	Confusion/disorientation	BUN $\geq 20$ mg/dL
	Platelets $< 100,000$	Hypotension requiring fluids
		WBC $\leq 4,000$ cells/mL
		Hypothermia

## Empiric Treatment



## Suggested Duration of Therapy

Most patients should be treated for a minimum of 5 days, should be afebrile for 48-72h, and should have no more than 1 of the following:

- Heart rate  $> 100$  beats/min
- Respiratory rate  $> 24$  breaths/min
- Systolic blood pressure  $< 90$  mmHg
- Altered mental status
- Arterial O<sub>2</sub> saturation  $< 90\%$

Patients with confirmed *MRSA* or *PSA* infection should be treated for a minimum of 7 days. In general, patients are treated for 1-2 past clinical stability.

# HOSPITAL & VENTILATOR ASSOCIATED PNEUMONIA

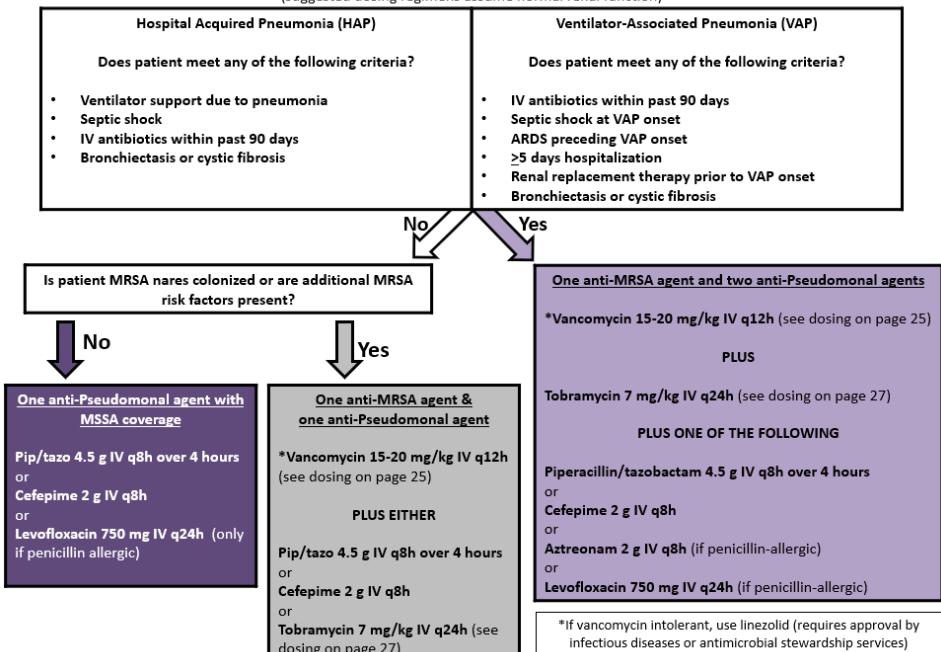
6

## Definitions

- **HAP (hospital acquired pneumonia):** Pneumonia that occurs >48 hours after admission that was not incubating at the time of admission and is not associated with mechanical ventilation
- **VAP (ventilator associated pneumonia):** Pneumonia that occurs >48 hours after endotracheal intubation

## Recommended Empiric Therapy (listed in order of preference)

(suggested dosing regimens assume normal renal function)



## Suggested Duration of Therapy

- 7 day treatment course
- Shorter or longer treatment durations may be indicated depending on the causative organism and rate of overall improvement

## Clinical Pearls

- Antimicrobial therapy should be de-escalated once culture results are available
- Consider a respiratory viral panel to help rule out viral respiratory infection

# DIABETIC FOOT INFECTIONS

7

## Diagnosis

Consider the possibility of infection occurring in any foot wound in a patient with diabetes

Infection present (2+ of the following criteria)

- Local swelling or induration
- Erythema (>0.5 cm)
- Local tenderness or pain
- Local warmth
- Purulent discharge

Plain radiographs of the affected foot to look for bony abnormalities (deformity, destruction) as well as soft tissue gas/foreign bodies is recommended for ALL patients

## Classification

Clinical Manifestations of Infection	PEDIS Grade	IDSA Infection Severity
No signs or symptoms of infection	1	Uninfected
Local infection involving ONLY the skin and subcutaneous tissue If erythema present, must be >0.5 cm and <2cm around the ulcer Exclusion of other causes of inflammatory response of the skin (trauma, gout, fracture, etc)	2	Mild
Local infection with erythema >2cm OR involving structures deeper than skin and subcutaneous tissues No systemic inflammatory response signs (SIRS)	3	Moderate
Local infection with signs of SIRS as manifested by 2+ of the following criteria Temperature >38C or <36C HR >90 beats/minute RR >20 breaths/minute or PaCO <sub>2</sub> <32 mmHg WBC >12,000 or <4000 cells/mm <sup>3</sup> or >10% bands	4	Severe

## Major Causative Organisms

Aerobes	Anaerobes
<i>Staphylococcus aureus</i>	<i>Bacteroides spp</i>
Coagulase-negative staphylococci	<i>Peptostreptococcus spp</i>
Beta-hemolytic streptococci	<i>Peptococcus spp</i>
<i>Enterococcus spp</i>	<i>Finegoldia magna</i>
<i>Enterobacteriaceae</i>	
<i>Proteus</i> , <i>Klebsiella</i> , & <i>Enterobacter spp</i>	
<i>Pseudomonas aeruginosa</i>	

## Empiric Antimicrobial Treatment (in addition to wound care/debridement)

\*\* Definitive treatment should be based off culture and sensitivity testing of the wound specimen \*\*

Uninfected wound	No antibiotic therapy necessary
Mild-moderate infection	
• No recent antimicrobial use	Cephalexin or amoxicillin/clavulanate
• Recent antimicrobial use (within 1 month)	PO: Levofloxacin IV: Ampicillin/sulbactam or levofloxacin
• Concern for MRSA	PO: Doxycycline or SMX/TMP or clindamycin IV: Vancomycin
Severe infection	Vancomycin PLUS Piperacillin/tazobactam or ceftazidime (consider addition of metronidazole if using cephalosporin)

\*\* If evidence of subcutaneous gas/emphysema on imaging, add clindamycin for anti-toxin effects \*\*

## When to empirically cover *Pseudomonas aeruginosa*?

- Presence of risk factors including high local prevalence of PSA infection and frequent exposure of the foot to water

## When to empirically cover MRSA?

- Patient is colonized with MRSA or has a history of MRSA infection within the past year
- The infection is sufficiently severe that failing to empirically cover MRSA would pose an unacceptable risk of treatment failure

# C. DIFFICILE INFECTION

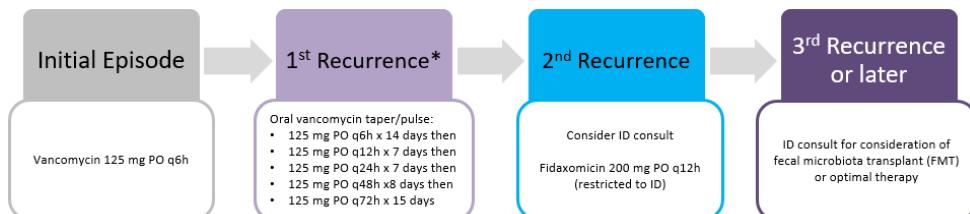
8

## Diagnosis

>3 unformed stools in 24 (or fewer) consecutive hours AND either a positive stool test for *C. difficile* or colonoscopic/histopathologic findings of pseudomembranous colitis

## Recommended Management

Clinical Definition	Supportive Clinical Data	Empiric Treatment
Asymptomatic carriage	Positive <i>C. difficile</i> PCR <b>WITHOUT</b> diarrhea, ileus, or colitis	No treatment necessary
Non-severe	Leukocytosis <15,000 cells/ $\mu$ g <b>AND</b> SCr <1.5 mg/dL	Use algorithm below
Severe	Leukocytosis $\geq$ 15,000 cells/ $\mu$ g <b>OR</b> SCr $\geq$ 1.5 mg/dL	Use algorithm below
Fulminant	Hypotension or shock, ileus, megacolon, or perforation	<b>Vancomycin 500 mg PO/NG q6h PLUS metronidazole 500 mg IV q8h</b> <ul style="list-style-type: none"><li>• Consider adding vancomycin PR if ileus present</li><li>• Recommend surgery consult</li></ul>



\*Available via order set

Orders tab → Order sets → Medicine orders → 6. Disease specific → Clostridium Difficile Infection → First recurrence

## Suggested Treatment Duration

- 10 day treatment course
- Consider extension to 14 days if patient has improved but not had symptom resolution at 10 days

## Clinical Pearls

- Stop offending agents when possible (systemic antibiotics, PPIs, H2 antagonists, etc.)
- In general, avoid the use of anti-motility agents, especially before treatment has started
- Use soap and water instead of EtOH based hand sanitizers
- Do NOT repeat stool test for a test of cure. This test may remain positive for weeks after treatment
- Certain antibiotics can predispose to a patient to *C. diff* infection. Help prevent infection by avoiding these antibiotics (fluoroquinolones, clindamycin, and broad-spectrum penicillins and cephalosporins) when appropriate and using antibiotics for the shortest effective duration
- Stress environmental cleaning and disinfection to prevent re-infection
  - Patients should wash hands with soap and water after using the bathroom, before eating or food preparation, and when hands are visibly soiled.
  - Patients with diarrhea should avoid using the same toilet as other family members.
  - Bathroom and kitchen areas (including toilet seats, toilet bowl, flush handle, sink faucet handles, and countertops) should be cleaned with a mixture of bleach and water (1-part bleach to every 10 parts water) to help prevent spread of infection.

# INFECTIVE ENDOCARDITIS

9-10

## Diagnosis

Based on modified Duke's criteria

Definite endocarditis: Presence of 2 major OR  
1 major and 3 minor OR  
5 minor criteria

Possible endocarditis Presence of 1 major and 1 minor OR  
3 minor criteria

<i>Major Criteria</i>	<i>Minor Criteria</i>
1) Two separate blood cultures positive for a typical microorganism causing IE 2) Evidence of endocardial involvement	1) Predisposing conditions 2) Fever >38°C 3) Embolic event 4) Immunologic phenomenon 5) Positive blood cultures that do not meet major criteria guidelines

## Major Criteria

1. Blood culture positive for IE
  - Typical microorganisms consistent with IE from 2 separate blood cultures (*Viridans Streptococci, Strep. gallolyticus/bovis, HACEK group, Staph. aureus or CA-enterococci*) in the absence of a primary focus
  - Microorganisms consistent with IE from persistently positive blood cultures defined as follows: At least 2 positive cultures of blood samples drawn >12h apart or all of 3 or a majority of >4 separate cultures of blood (first and last sample drawn at least 1 hour apart)
  - Single positive blood culture for *Coxiella burnetii* or anti-phase 1 IgG antibody titer >1:800
2. Evidence of endocardial involvement
  - Echocardiogram positive for IE; oscillating intracardiac mass on valve or supporting structures, in the path of regurgitant jets, or on implanted material in the absence of an alternative anatomic explanation; new abscess; new partial dehiscence of prosthetic valve; new valvular regurgitation

## Minor Criteria

1. Predisposing conditions
2. Fever >38 °C
3. Embolic event or vascular phenomena
  - Major arterial emboli, septic pulmonary infarcts, mycotic aneurysm, intracranial hemorrhage, conjunctival hemorrhage, Janeway lesions
4. Immunologic phenomena
  - Glomerulonephritis, Osler's nodes, Roth's spots, rheumatoid factor
5. Positive blood cultures that do not meet major criteria guidelines

## Suggested Treatment Regimens/Duration by Causative Organism

See tables to the right

### **Streptococcal species**

<b>Native valve</b>		
MIC (mcg/mL)	First-line therapy	Duration of treatment
≤0.12 (age >65 or poor renal function)	Penicillin G 12-18 million units/24h OR ceftriaxone 2 g q24h	4 weeks
≤0.12 (good renal function)	Penicillin G or ceftriaxone (above doses) AND gentamicin 1.5 mg/kg q12h	2 weeks 2 weeks
>0.12 and ≤0.5	Penicillin G 24 million units/24h OR ceftriaxone 2 g q24h AND gentamicin 1.5 mg/kg q12h	4 weeks 2 weeks
<b>Prosthetic valve</b>		
MIC (mcg/mL)	First-line therapy	Duration of treatment
≤0.12	Penicillin G 24 million units/24h OR ceftriaxone 2 g q24h AND gentamicin 1.5 mg/kg q12h	6 weeks 2 weeks
>0.12	Penicillin G 24 million units/24h OR ceftriaxone 2 g q24h AND gentamicin 1.5 mg/kg q12h	6 weeks 6 weeks

### **Staphylococcal species**

<b>Native valve</b>		
Etiologic organism	First-line therapy	Duration of treatment
MSSA or MSSE	Nafcillin 2 g q4h OR ceftazolin 2 g IV q8h	6 weeks
MRSA or MRSE	Vancomycin 15 mg/kg q12h (goal trough 15-20 mcg/mL)	6 weeks
<b>Prosthetic valve</b>		
Etiologic organism	First-line therapy	Duration of treatment
MSSA or MSSE	Nafcillin 2 g q4h AND rifampin 300 mg PO q8h AND gentamicin 1.5 mg/kg IV q12h	≥6 weeks ≥6 weeks 2 weeks
MRSA or MRSE	Vancomycin 15 mg/kg q12h (goal trough 15-20 mcg/mL) AND rifampin 300 mg PO q8h AND gentamicin 1.5 mg/kg q12h	≥6 weeks ≥6 weeks 2 weeks

### **Enterococcal species**

<b>Native or prosthetic valve</b>		
Etiologic organism	First-line therapy	Duration of treatment
<i>Enterococci</i> susceptible to penicillin, gentamicin	Ampicillin 2 g q4h OR penicillin G 18-30 million units/24h AND gentamicin 1.5 mg/kg q12h  OR Ampicillin 2 g q4h AND ceftriaxone 2 g q12h	4-6 weeks 4-6 weeks  6 weeks
<i>Enterococci</i> resistant to penicillin	Vancomycin 15 mg/kg q12h (goal trough 15-20 mcg/mL) AND gentamicin 1.5 mg/kg q12h	6 weeks 6 weeks
<i>Enterococci</i> resistant to gentamicin	Ampicillin 2 g q4h AND ceftriaxone 2 g q12h	6 weeks
<i>Enterococci</i> resistant to vancomycin	Daptomycin 10-12 mg/kg q24h OR linezolid 600 mg IV/PO q12h	>6 weeks
HACEK organisms	Ceftriaxone 2 g q24h OR ampicillin 2 g q4h OR ciprofloxacin 400 mg IV q12h	4 weeks

HACEK organisms:

*Haemophilus spp*, *Actinobacillus actinomycetemcomitans*, *Cardiobacterium hominis*, *Eikenella corrodens*, *Kingella kingae*

# NEUTROPENIC PROPHYLAXIS

11

Suggested Antimicrobial Prophylaxis	Bacterial	Fungal	Viral	PCP
<b>Low Risk</b> <ul style="list-style-type: none"> <li>Anticipated duration of neutropenia &lt;7 days</li> <li>Standard chemotherapy regimens for most solid tumors</li> </ul>			✓*	
<b>Intermediate Risk</b> <ul style="list-style-type: none"> <li>Anticipated duration of neutropenia 7-10 days</li> <li>CLL, multiple myeloma, lymphoma, autologous HCT</li> <li>Purine analog therapy (fludarabine, cladribine, etc.)</li> </ul>	✓	✓	✓	✓
<b>High Risk</b> <ul style="list-style-type: none"> <li>Anticipated duration of neutropenia &gt;10 days</li> <li>Leukemia, allogeneic HCT</li> <li>Alemtuzumab therapy</li> <li>GVHD treated with high dose steroids (&gt;20 mg prednisone or equivalent per day)</li> </ul>	✓	✓	✓	✓

\*Antiviral prophylaxis indicated only if patient had prior HSV episode

Antimicrobial Prophylaxis	
<ul style="list-style-type: none"> <li>levofloxacin 750 mg PO q24h or</li> <li>ciprofloxacin 500-750 mg PO q12h and amoxicillin/clavulanate 875/125 mg PO q12h</li> </ul>	
Antifungal Prophylaxis	
Disease (intermediate to high risk)	Suggested First-Line Prophylaxis
ALL	fluconazole 400 mg PO q24h
MDS, AML (neutropenic)	<ul style="list-style-type: none"> <li>posaconazole 300 mg PO q12h x2 doses then 300 mg PO q24h (restricted to ID) or</li> <li>voriconazole 400 mg PO q12h x2 doses then 200 mg PO q12h (restricted to ID)</li> </ul>
Autologous HCT with mucositis	fluconazole 400 mg PO q24h
Autologous HCT without mucositis	consider no prophylaxis
Allogeneic HCT (neutropenic)	fluconazole 400 mg PO q24h
Significant GVHD	<ul style="list-style-type: none"> <li>posaconazole 300 mg PO q12h x2 doses then 300 mg PO q24h (restricted to ID) or</li> <li>voriconazole 400 mg PO q12h x2 doses then 200 mg PO q12h (restricted to ID)</li> </ul>
Other	fluconazole 400 mg PO QD
Antiviral Prophylaxis	Antipneumocystis Prophylaxis
Preferred:	Preferred:
<ul style="list-style-type: none"> <li>valacyclovir 500 mg PO q12h or q8h</li> <li>acyclovir 400 – 800 mg PO BID</li> </ul>	<ul style="list-style-type: none"> <li>TMP/SMZ 1 DS tablet PO q24h or</li> <li>TMP/SMZ 1 DS tablet PO 3x/week</li> </ul>
Alternative (restricted to ID):	Alternative (restricted to ID):
<ul style="list-style-type: none"> <li>famciclovir 250 mg PO BID or</li> </ul>	<ul style="list-style-type: none"> <li>atovaquone 1500 mg PO q24h with food</li> </ul>

# FEBRILE NEUTROPENIA

12

## Definitions

- Fever:** A single oral temperature  $>38.3^{\circ}\text{C}$  ( $101^{\circ}\text{F}$ ) or a temperature of  $>38.0^{\circ}\text{C}$  ( $100.4^{\circ}\text{F}$ ) sustained over a one-hour period
- Neutropenia:** ANC  $<500 \text{ cells/mm}^3$  or an ANC that is expected to decrease to  $<500 \text{ cells/mm}^3$  during the next 48 hours

## Common Etiologic Agents

Gram-positive organisms	Gram-negative organisms
Coagulase-negative staphylococci	<i>Escherichia coli</i>
<i>Staphylococcus aureus</i> (including MRSA)	<i>Klebsiella species</i>
<i>Enterococcus</i> species (including VRE)	<i>Enterobacter species</i>
Viridans group streptococci	<i>Pseudomonas aeruginosa</i>
<i>Streptococcus pneumoniae</i>	<i>Citrobacter species</i>
<i>Streptococcus pyogenes</i>	<i>Acinetobacter species</i>
	<i>Stenotrophomonas maltophilia</i>

## Anticipated Duration of Neutropenia

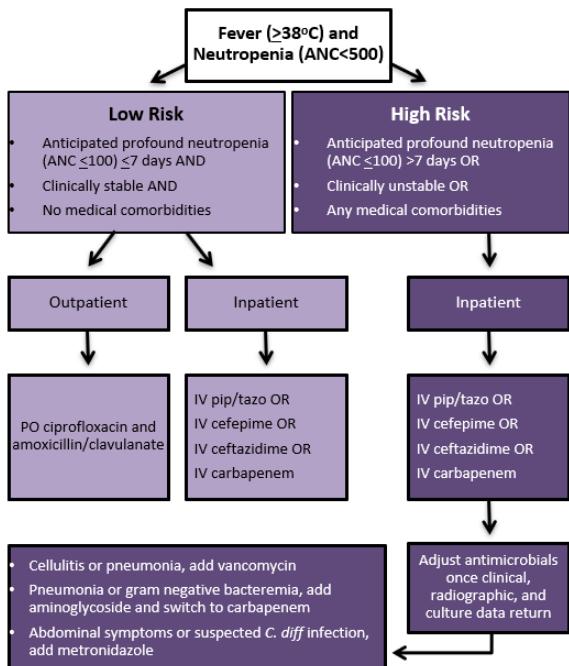
Refer to NCCN guidelines on the Prevention and Treatment of Cancer Related Infections

## Empiric Treatment

Use algorithm to the right

## Clinical Pearls

- Vancomycin is NOT recommended as a standard part of the initial empiric antibiotic regimen. Use vancomycin if:
  - Hemodynamic instability/evidence of severe sepsis
  - Pneumonia documented by imaging
  - Positive blood cultures for gram+ bacteria before final ID/susceptibilities
  - Suspected catheter-related infection
  - SSTI at any site
  - Colonization with MRSA
- Antibiotic prophylaxis with levofloxacin is recommended for high-risk patients expected to have prolonged and profound neutropenia
- Empiric antifungal therapy should be considered if persistent/recurrent fever after 4-7 days of antibiotics
- Antiviral therapy indicated for patients with documented HSV or VZV infection



Freifeld AG, Bow EJ, Sepkowitz KA, et al. Clinical Practice Guideline for the Use of Antimicrobial Agents in Neutropenic Patients with Cancer: 2010 Update by the Infectious Diseases Society of America. *Clin Infect Dis* 2011;52(4):e56–e93

National Comprehensive Cancer Network. Prevention and treatment of cancer related infections (Version 1.2016). [http://www.nccn.org/professionals/physician\\_gls/pdf/infections.pdf](http://www.nccn.org/professionals/physician_gls/pdf/infections.pdf) Accessed April 2016

# BACTERIAL MENINGITIS

13

## Diagnosis

- Clinical findings on lumbar puncture and CSF analysis
- Classic triad of fever, nuchal rigidity, and altered mental status
- Additional symptoms may include chills, vomiting, photophobia, severe headache, etc.

## Recommended Empiric Management

Patient Characteristics	Common Etiologic Agents	Empiric Treatment
Immunocompetent AND age < 50	<i>Streptococcus pneumoniae</i> <i>Neisseria meningitidis</i> <i>Haemophilus influenzae</i>	Ceftriaxone 2g IV q12h AND Vancomycin 30-45 mg/kg/day divided  *Add dexamethasone 0.15 mg/kg IV q6h x2-4 days if pneumococcal meningitis suspected
Immunocompetent AND age ≥ 50	<i>Streptococcus pneumoniae</i> <i>Listeria monocytogenes</i>	Ceftriaxone 2g IV q12h AND
Immunocompromised	<i>Neisseria meningitidis</i> <i>Haemophilus influenzae</i> Group B Streptococci	Vancomycin 30-45 mg/kg/day divided AND Ampicillin 2g IV q4h
Post neurosurgery, CSF shunt infection, or head trauma	<i>Staphylococcus aureus</i> Coagulase negative <i>Staphylococci</i> Aerobic gram-negative rods <i>Pseudomonas aeruginosa</i>	Vancomycin 30-45 mg/kg/day divided AND Cefepime 2g IV q8h
Severe penicillin allergy		Consult allergy service for desensitization  If Listeria suspected, add TMP/SMZ 5mg/kg IV q6h

## Suggested Duration of Therapy

Etiologic Agent	Recommended Treatment Duration
<i>Neisseria meningitidis</i>	7 days
<i>Haemophilus influenzae</i>	7 days
<i>Staphylococcus aureus</i>	10 -14 days
<i>Streptococcus pneumoniae</i>	10-14 days
Coagulase negative <i>Staphylococci</i> or <i>P. acnes</i>	10 – 14 days
<i>Streptococcus agalactiae</i>	14-21 days
Aerobic gram-negative rods	21 days
<i>Listeria monocytogenes</i>	>21 days

## Clinical Pearls

- Start antibiotics as soon as meningitis is suspected and optimize doses for CNS penetration
- Patient with any of the following criteria should receive a CT scan before lumbar puncture:
  - Immunocompromised
  - History of CNS disease
  - New onset seizure
  - Papilledema
  - Abnormal level of consciousness
  - Focal neurologic deficit
- If using dexamethasone, give 10-20 minutes before or concomitant with the first dose of antibiotics
- If a lumbar puncture must be delayed/deferred, obtain blood cultures and start empiric treatment

Tunkel AR, Hartman BJ, Kaplan SL et al. Practice Guidelines for the Management of Bacterial Meningitis. *Clin Infect Dis.* 2004; 39:1267-84

Tunkel AR, Hasbun R, Bhimraj AB et al. 2017 Infectious Diseases Society of America's Clinical Practice Guidelines for Healthcare-Associated Ventriculitis and Meningitis. *Clin Infect Dis.* 2017; 64(6):e34-65

# SKIN AND SOFT TISSUE INFECTIONS

14

## Definitions:

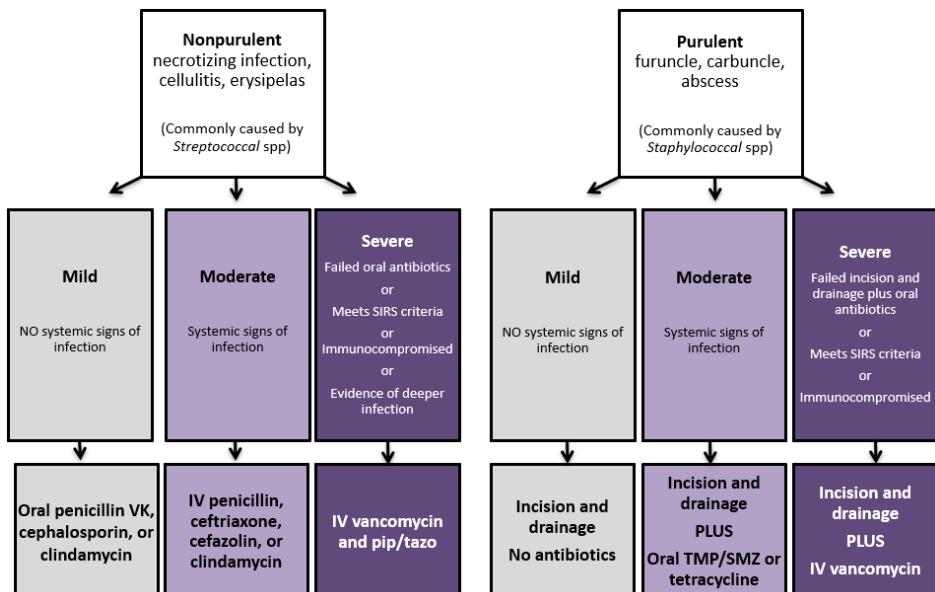
- **Furuncle:** Infection of the hair follicle also known as boils
- **Carbuncle:** Infection involving several adjacent furuncles
- **Abscess:** Collection of pus within the dermis and deeper skin tissues
- **Erysipelas:** Infection limited to the upper dermis, including the superficial lymphatics
- **Cellulitis:** Infection which involves the deeper dermis and subcutaneous fat
- **Necrotizing infection:** Deep infection which involves the fascial and/or muscle compartments and are potentially devastating due to major tissue destruction and death

## Empiric Treatment

- See flow chart below

## Recommended Duration of Therapy

Classification	Treatment Duration
Furuncles/carbuncles	No treatment usually required; self-limiting
Recurrent abscesses	5 – 10 days
Erysipelas/cellulitis	5 days; treatment should be extended if the infection has not improved within this time period
Recurrent cellulitis	Consider prophylactic antibiotics for 4–52 weeks
Necrotizing infections	Antimicrobial therapy should be administered until further debridement is no longer necessary, the patient has improved clinically, and fever has been absent for 48–72 hours



# INFLUENZA TREATMENT

15

## Diagnosis

- Based on the clinical presentation of the patient and the results of diagnostic testing
- History of influenza vaccination does NOT preclude influenza infection when signs and symptoms are compatible with the clinical syndrome

## Clinical Presentation

Common symptoms of influenza include abrupt onset of fever, chills, myalgias, headache, and/or fatigue

Signs and Symptoms	Influenza	Cold
Symptom onset	Abrupt	Gradual
Fever	Usual, lasts 3-4 days	Rare
Aches	Usual, often severe	Slight
Chills	Fairly common	Uncommon
Fatigue, weakness	Usual	Sometimes
Sneezing	Sometimes	Common
Stuffy nose	Sometimes	Common
Sore throat	Sometimes	Common
Chest discomfort/cough	Common; can be severe	Mild to moderate
Headache	Common	Rare

## Treatment

- Decision to treat should not wait for the confirmation of laboratory results
- Treatment should be continued in the setting of negative diagnostic results in patients with signs/symptoms suggestive of influenza infection. False negative results are possible

Antiviral treatment should begin within 48 hours of symptom onset

## Recommended Treatment (based on renal function)

Treatment regimen	Estimated Creatinine Clearance (ml/min)
Oseltamivir 75 mg PO BID x 5 days	>60
Oseltamivir 30 mg PO BID x 5 days	31-60
Oseltamivir 30 mg PO daily x 5 days	11-30
Oseltamivir 30 mg immediately and then 30 mg after every hemodialysis session for 5 days (assuming three HD sessions in the 5-day period)	ESRD on hemodialysis
Oseltamivir 30 mg PO once	Peritoneal dialysis

# INFLUENZA PREVENTION

The CDC recommends a yearly flu vaccine as the first and most important step in protecting against influenza

Post-exposure prophylaxis is an option for select patients who meet ALL three of the following criteria:

- High-risk for influenza complications (any of the following)
  - Age ≥ 65 years old
  - Immunosuppression caused by medication or HIV infection
  - Pregnant women (and women up to two weeks postpartum)
  - Chronic health conditions including chronic pulmonary, cardiovascular (except hypertension alone), renal, hepatic, hematologic (including sickle cell disease), metabolic disorders (including diabetes mellitus), neurologic conditions (disorders of the brain, spinal cord, nerve, muscle, epilepsy, stroke, or intellectual disability)
- Did NOT receive this year's influenza vaccine
- Present within 48 hours of close contact with an infected individual

## Recommended Post-Exposure Prophylaxis (based on renal function)

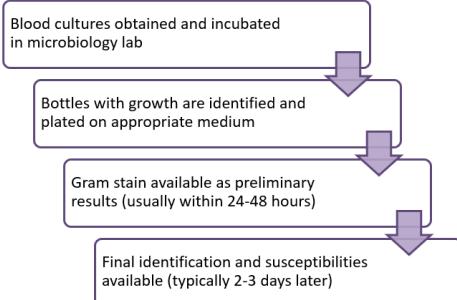
Regimen	Estimated Creatinine Clearance (ml/min)
Oseltamivir 75 mg PO daily x 10 days	>60
Oseltamivir 30 mg PO daily x 10 days	31-60
Oseltamivir 30 mg PO every other day x 10 days	11-30
Oseltamivir 30 mg immediately and then 30 mg after every other hemodialysis sessions for the recommended prophylaxis duration	ESRD on hemodialysis
Oseltamivir 30 mg immediately and additional 30 mg given 7 days later	Peritoneal dialysis

# BACTEREMIA

16

## Blood Culture Basics

- 1 blood culture = 2 bottles (1 aerobic and 1 anaerobic bottle)
- Typically order two sets of blood cultures from different anatomical sites = total of four bottles
- Blood cultures take 5 days to finalize
- Growth in anaerobic bottle does NOT necessarily mean you're growing an anaerobe
- Use the guide to Gram-stain and morphology on page #17 to guide treatment choices before final identification and susceptibilities are available



Gram stain results	Likely organism (by species)	Contaminant?
Gram positive cocci in clusters	<ul style="list-style-type: none"><li>• <i>Staphylococcus</i></li></ul>	Possible; follow up final identification <ul style="list-style-type: none"><li>• <i>S. aureus/lugdunensis</i> = Requires treatment</li><li>• Coagulase negative Staphylococci = May not require treatment</li></ul>
Gram-positive cocci in chains/pairs	<ul style="list-style-type: none"><li>• <i>Streptococcus</i></li><li>• <i>Enterococcus</i></li></ul>	Likely not
Gram-positive rods	<ul style="list-style-type: none"><li>• <i>Bacillus</i></li><li>• <i>Clostridium</i></li><li>• <i>Listeria</i></li><li>• <i>Corynebacterium</i></li><li>• <i>Propionibacterium</i></li></ul>	Possible; follow up final identification <ul style="list-style-type: none"><li>• <i>Bacillus anthracis</i>, <i>Listeria monocytogenes</i>, or <i>Clostridium spp</i> = Requires treatment</li><li>• <i>Bacillus cereus/non-anthrax</i>, <i>Corynebacterium spp.</i>, or <i>Propionibacterium acnes</i> = May not require treatment</li></ul>
Gram-negative rods	<ul style="list-style-type: none"><li>• <i>Enterobacteriaceae</i> family</li><li>• SPACE/SPICE organisms</li></ul>	Likely not
Yeast	<ul style="list-style-type: none"><li>• <i>Candida</i></li><li>• <i>Cryptococcus</i></li></ul>	Likely not

***Staphylococcus aureus* in blood cultures is NEVER a contaminant. Formal ID consult is recommended for classification, additional workup, and optimal treatment regimen/duration**



## Approach to Possible Contaminants

Use clinical judgement

- Is the patient hemodynamically unstable or septic?
- Does the preliminary culture suggest an organism that is known to cause endocarditis?
- How many bottles are positive?

Weigh risk vs benefit

- Risk of not treating "true" bacteremia
- Risk of inappropriate antimicrobial therapy

Call the Infectious Diseases or Antimicrobial Stewardship team for assistance in interpreting positive blood cultures

Factors Suggesting Contaminant	Factors Suggesting True Bacteremia
<ul style="list-style-type: none"><li>• Extended time to positivity</li><li>• Certain bacterial strains</li><li>• Multiple &gt; single pathogen</li></ul>	<ul style="list-style-type: none"><li>• Single &gt; multiple pathogens</li><li>• Multiple BCx with the same organism</li><li>• Typical pathogens</li><li>• Clinical signs of infection</li><li>• Presence of risk factors</li></ul>

## Other Considerations

- Any patient with bacteria in the blood (contaminant or real) should get 2 sets of repeat blood cultures to document clearance. Duration of therapy typically starts from the date of the first negative blood culture
- Blood cultures growing certain organisms may need central line removal and/or work-up to rule out endocarditis
- Use bactericidal antibiotics for treatment
- Avoid PO antibiotics (with the exception of fluoroquinolones for select Gram negative bacteremia) given issues with poor bioavailability and serum concentrations at the site of infection

# CHOOSING AN ANTIBIOTIC

17

## What is Minimum Inhibitory Concentration (MIC)?

- Minimum inhibitory concentration is the lowest concentration of an antimicrobial that *inhibits* bacterial growth
- The MIC ranges are different for each antibiotic tested and different for each organism

**MICs are NOT directly comparable. Do not simply choose the antibiotic with the lowest MIC.**

- Comparing MIC values between antibiotics is similar to comparing "apples to oranges". For example, a ciprofloxacin MIC of 2 is not better than a piperacillin/tazobactam MIC of 8

## Guide to Interpreting a Culture and Susceptibility Report



### R (resistant)

Choose an alternative agent

### I (intermediate)

If no other options available, contact infectious disease specialist to see if resistance can be overcome with high doses

### S (susceptible)

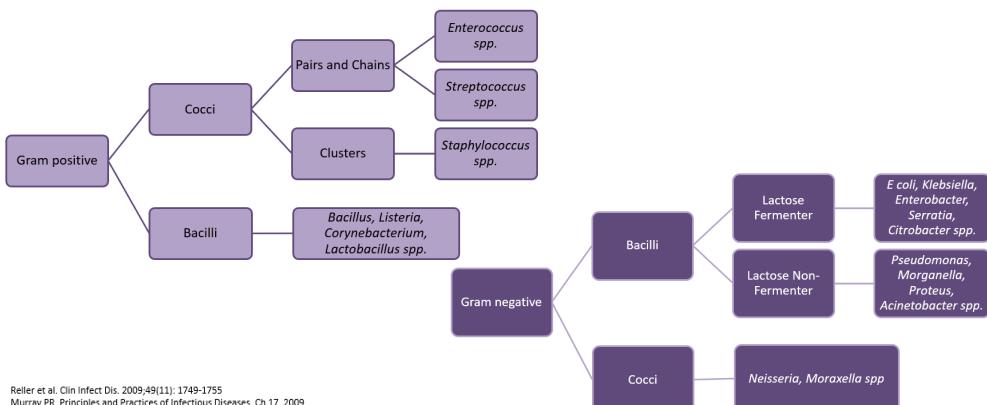
Good option in-vitro; examine patient-specific factors

- 1) Pick the ideal antibiotic for specific situation
- 2) If that antibiotic is susceptible, consider your patient and whether that choice is appropriate:
  - Drug allergies?
  - Renal function?
  - Liver function?
  - Drug interactions?
  - Site of action?

- Keep in mind in vitro susceptibility does not guarantee in vivo efficacy. There are MANY scenarios where an antimicrobial may appear susceptible but would be a poor choice for treatment.
- The MIC values do NOT suggest chance of clinical success. Clinical success is dependent upon:
  - Host responses
  - Site of infection
  - Bacterial toxin production
  - Presence of biofilms
  - Medication pharmacodynamics
  - Other factors
- Susceptibility to a drug within a class does not always correspond to susceptibility to all drugs in that class
- Susceptibility tests are assessed based on plasma levels and may not predict concentrations at the site of infection. They do not take into account local factors such as pus, necrosis, poor perfusion, etc

## Guide to Gram Stains and Morphology

- Preliminary culture data is reported as a Gram stain.
- Gram stains can help guide treatment decisions before an organism identification/susceptibilities are known
- The antibiogram (on pages 29-30 & the intranet) can assist you in choosing an appropriate antibiotic



# RENAL DOSING GUIDE

18

## Abbreviations

ABW:	Actual body weight	GFR:	Glomerular filtration rate
AdjBW:	Adjusted body weight	IBW:	Ideal body weight
CrCl:	Creatinine clearance	SCr:	Serum creatinine

## Estimating Renal Function Using the Cockcroft Gault Equation

- If the patient's ABW < IBW, use ABW
- If the patient's ABW > 1.3x IBW, use AdjBW

### Ideal Body Weight

Men:	50 kg + (2.3 kg per inch over 5 feet)
Women:	45 kg + (2.3 kg per inch over 5 feet)

### Adjusted Body Weight

$$\text{AdjBW} = 0.4 (\text{ABW} - \text{IBW}) + \text{IBW}$$

### Cockcroft Gault Equation

$$\text{CrCl (mL/min)} = \frac{(140 - \text{age}) (\text{weight in kg}) (0.85 \text{ if female})}{72 \times \text{SCr}}$$

### Cystatin-C

- Better estimation of renal function in spinal cord injury patients with low muscle mass

Cystatin-C	eGFR
1.0	75-100
1.1	68-86
1.2	62-75
1.3	55-65
1.4	50-58
1.5	46-52
1.6	42-47
1.7	40-43
1.8	37-39
1.9	34-36

Cystatin-C	eGFR
2.0	32-34
2.1	30-32
2.2	28-30
2.3	26-28
2.4	25-27
2.5	24-26
2.6	22-24
2.7	21-23
2.8	20-22
2.9	18-21

Cystatin-C	eGFR
3.0	17-20
3.1	17-20
3.2	16-19
3.3	15-17
3.4	14-16
3.5	14-16
3.6	13-16
3.7	12-15
3.8	12-15
3.9	11-15

## Antimicrobials Typically Not Requiring Renal Dosage Adjustment

Antibiotics		
Azithromycin	Fidaxomicin	Moxifloxacin
Cefuroxime	Fosfomycin	Nafcillin
Ceftriaxone	Isoniazid	Penicillin VK (oral only)
Clindamycin	Linezolid	Rifampin
Doxycycline	Metronidazole	Tigecycline
Erythromycin	Minocycline	Vancomycin (oral only)
Antifungals		
Amphotericin B	Ketoconazole	Posaconazole (oral only)
Isavuconazonium sulfate	Micafungin	Voriconazole (oral only)
Antivirals/Antiretrovirals		
Abacavir	Dolutegravir <sup>b</sup>	Lopinavir/ritonavir
Atazanavir	Efavirenz	Raltegravir
Daclatasvir	Elbasvir/grazoprevir	Rilpivirine
Darunavir	Etravirine	Ritonavir

<sup>a</sup>No dosage adjustments in treatment-naïve or treatment-experienced/INSTI-naïve patients

# DOSING GUIDELINES BASED ON RENAL FUNCTION: IV AGENTS

19

Antimicrobial	Typical Dose (may vary)	CrCl (ml/min)	Suggested Dose Adjustment
Acyclovir	5-10 mg/kg q8h	25-50 10-24 <10	5-10 mg/kg q12h 5-10 mg/kg q24h 2.5-5 mg/kg q24h
*Amikacin	See aminoglycoside dosing/monitoring guide on pages 26-28		
Ampicillin	1-2 g q4-6h	10-50 <10	1-2 g q6-12h 1-2 g q12-24h
Ampicillin/sulbactam	1.5-3 g q6h	15-29 <15	1.5-3 g q12h 1.5-3 g q24h
Aztreonam	2 g q8h	10-29 <10	1 g q8h 500 mg q8h
Cefazolin	1-2 g q8h	11-34 <11	0.5-1 g q12h 0.5-1 g q24h
Cefepime	See alternative dosing on page 23		
Cefotaxime	1-2 g q6-8h	<20	0.5-1 g q6-8h
Cefotetan	1-2 g q12h	10-30 <10	0.5-1 g q12h 250-500 mg q12h
Cefoxitin	1-2 g q6-8h	30-50 10-29 5-9	1-2 g q8-12h 1-2 g q12-24h 0.5-1 g q12-24h
*Ceftaroline	600 mg q12h	31-50 15-30 <15	400 mg q12h 300 mg q12h 200 mg q12h
Ceftazidime	1-2 g q8h	31-50 16-30 ≤15	1-2 g q12h 1-2 g q24h 0.5-1 g q24h
*Ceftazidime/avibactam	2.5 g q8h	31-50 16-30 6-15 ≤5	1.25 g q8h 0.94 g q12h 0.94 g q24h 0.94 g q48h
*Ceftolozane/tazobactam	1.5 g q8h (most indications)	30-50 15-29	750 mg q8h 375 mg q8h
	3 g q8h (PAP/VAP or serious multi-drug resistant <i>Pseudomonas</i> infections)	30-50 15-29	1.5 g q8h 750 mg q8h
Cefuroxime	1.5 g q8h	10-20 <10	1.5 g q12h 1.5 g q24h
Ciprofloxacin	400 mg q12h (most indications) 400 mg q8h (anti- <i>Pseudomonas</i> dosing)	<30 <30	400 mg q24h 400 mg q12h
*Colistimethate	Consult ID PharmD		
*Daptomycin	4-12 mg/kg q24h	<30	4-12 mg/kg q48h
*Ertapenem	1 g q24h	<30	500 mg q24h
Fluconazole	200-800 mg q24h	<50	Same LD → 100-400 mg q24h
Gentamicin	See aminoglycoside dosing/monitoring guide on pages 26-28		
*Imipenem/cilastatin	1 g q8h	60-89 30-59 15-29	500 mg q6h 500 mg q8h 500 mg q12h
Levofloxacin	500 mg q24h (most indications) 750 mg q24h (anti- <i>Pseudomonas</i> dosing)	20-49 <20 20-49 <20	500 mg LD → 250 mg q24h 500 mg LD → 250 mg q24h 750 mg q48h 750 mg LD → 500 mg q48h
*Meropenem	See alternative dosing on page 23		
*Meropenem/vaborbactam	4 g q8h	30-49 15-29 <15	2 g q8h 2 g q12h 1 g q12h
Penicillin G	2-4 million units q4h	10-50 <10	75% of the normal dose 20% to 50% of the normal dose
Piperacillin/tazobactam	See alternative dosing on page 23		
*Polymyxin B	Consult ID PharmD		
TMP/SMZ	8-20 mg/kg/TMP/day divided q6-12h	15-30 <15	Reduce dose to 50% usual dose Reduce dose 25-50% usual dose
*Tobramycin	See aminoglycoside dosing/monitoring guide on pages 26-28		
Vancomycin	See vancomycin dosing/monitoring guide on pages 25		
*Voriconazole	4 mg/kg q12h	<50	Avoid use, consider alternative or use PO

\* = Restricted to Infectious Diseases/Antimicrobial Stewardship

BOLD = Nonformulary; requires PADR consult approval

LD = loading dose

# DOSING IN DIALYSIS

20

Antimicrobial	Hemodialysis (HD) (Regimens assume three complete HD sessions per week. Dose after HD when applicable)	Peritoneal dialysis (IV doses given below. Contact PharmD for intraperitoneal dosing)	CVVHDF
Acyclovir	2.5 mg/kg q12h	2.5-5 mg/kg q24h	5-10 mg/kg q12-24h
* Amikacin	7.5 mg/kg LD → 5-7.5 mg/kg post-HD. Re-dose when pre-HD level <10mcg/mL	Consider 7.5 mg/kg LD then follow levels	10 mg/kg LD → 7.5 mg/kg q24-48h Re-dose when level <10 mcg/mL
Ampicillin	1-2 g q12-24h	1-2 g q12-24h	2 g LD → 1-2 g q6-8h
Ampicillin/Sulbactam	1.5-3 g q12-24h	1.5 g q12h or 3 g q24h	3 g q8-12h
Aztreonam	500 mg q12h	25% of usual dose at the usual interval	1 g q8h or 2 g q12h
Cefazolin	Daily: 1 g q24h 3x/week: 6 g/2 g/2 g or 2 g/2 g/3 g post-HD	500 mg q12h	1 g q8h or 2 g q12h
Cefepime	Daily: See dosing on page 23 3x/week: 2 g/2 g/2 g post-HD	1 g q24h	2 g q8-12h
Cefotaxime	1-2 g q24h	1 g q24h	1-2 g q6-8h
*Ceftaroline	200 mg q12h	No data	400-600 mg q12h
Ceftazidime	Daily: 1 g q24h 3x/week: 1 g/1 g/1 g or 1 g/1 g/2 g post-HD	1 g q24h	2 g q8-12h
*Ceftriaxone/Avibactam	0.94 g q24-48h based on eCrCl	No data	1.25 g q8h
*Ceftiofozane/tazobactam	750 mg LD → 150 mg q8h (normal dose 1.5g q8h) 2.25 g LD → 450 mg q8h (normal dose 3g q8h)	No data	3 g LD → 750 mg q8h
Ciprofloxacin	PO: 250-500 mg q24h IV: 200-400 mg q24h	PO: 250-500 mg q24h IV: 200-400 mg q24h	PO: N/A IV: 400 mg q12h
*Colistimethate	Contact ID PharmD	No data	Contact ID PharmD
*Daptomycin (based on 6 mg/kg dose)	Preferred: 6 mg/kg q24h Alternative: 6 mg/kg q18h	6 mg/kg q48h	6 mg/kg q24h
*Ertapenem	Daily: 500 mg q24h 3x/week: 1 g/1 g/1 g post-HD	500 mg q24h	1 g q24h
Filconazole (based on 400 mg dose)	800 mg LD → 400 mg 3x/week post-HD when pre-HD level <1 (synergy/mild infxn), <1.5-2 (moderate infxn), or <3.5 (severe infxn)	800 mg LD → 200 mg q24h Consider 2-3 mg/kg LD then follow levels	800 mg q24h 2-3 mg/kg LD → 1-2 mg/kg q24h. Re-dose when level <1 (synergy/mild infxn), <1.5-2 (moderate infxn), or <3.5 (severe infxn)
Gentamicin	2-3 mg/kg LD → 1-2 mg/kg post-HD. Re-dose when pre-HD level <1 (synergy/mild infxn), <1.5-2 (moderate infxn), or <3.5 (severe infxn)	500 mg LD → 250 mg q48h 750 mg LD → 500 mg q48h or 250 mg q24h	500 mg LD → 250 mg q24h 750 mg LD → 500 mg q24h
Levofloxacin (500 mg q24h)	500 mg LD → 250 mg q48h 750 mg LD → 500 mg q48h or 250 mg q24h	0.5-1 g q24h	0.5-1 g q24h
Levofloxacin (750 mg q24h)	See dosing on page 23	No data	No data
*Meropenem	1-2 g q12h based on eCrCl	No data	4 mu LD → 2-4 mu q4-6h
*Meropenem/vaborbactam	Typical LD → 25-50% normal dose q4-6h	See dosing on page 23	See dosing on page 23
Penicillin G (mu=million units)	See dosing on page 23	No data	Contact ID PharmD
Piperacillin/tazobactam	Contact ID PharmD	Same dosing as for gentamicin above	Same dosing as for Gentamicin above
Tobramycin	Same dosing as for gentamicin above	Reduce dose to 25-50% usual dose	No dose adjustment necessary
TMP/SMZ	25 mg/kg LD → 500-1000 mg or 5-10 mg/kg 3x/week post-HD	1000 mg q4-7 days	25 mg/kg LD → 1000 mg q24h or 7.5-10 mg/kg q12h
Vancomycin			

Other dosing regimens may be appropriate. Above regimens compiled from individual package inserts, Lexicomp®, expert opinion, Heinz et al *Pharmacotherapy* 2009;29(5):562-577, and Pistolese et al. *Antimicrob. Agents Chemother.* 2019;63(8):1-18

# DOSING GUIDELINES BASED ON RENAL FUNCTION: PO AGENTS

21

Antimicrobial	Typical Dose (may vary)	eCrCl (ml/min)	Suggested Dose Adjustment
Acyclovir	400-800 mg q12h (neutropenic ppx) All other indications → use valacyclovir	<10	200 mg q12h
*Adefovir	10 mg q24h	30-49 10-29	10 mg q48h 10 mg q72h
Amoxicillin	500 mg q8h (most indications) or 1000 mg q12h (part of <i>H pylori</i> regimen) 1000 mg q8h ( <i>Actinomycosis</i> )	10-29 <10 10-29 <10	500 mg q12h 500 mg q24h 1000 mg q12h 500 mg q12h
Amoxicillin/clavulanate	875 mg q12h or 500 mg q8h	<30 10-30 <10	Avoid 875 mg tablets 250-500 mg q12h 250-500 mg q24h
*Cefadroxil	500-1000 mg q12h	25-50 10-25	500 mg q12h 500 mg q24h
Cefdinir	300 mg q12h	<30	300 mg q24h
*Cefpodoxime	100-400 mg q12h	<30	100-400 mg q24h
Cephalexin	500 mg q6h	30-59 15-29 5-14	500 mg q12h 250 mg q8-12h 250 mg q24h
Ciprofloxacin (IR)	500 mg q12h (most indications) or 750 mg q12h (anti- <i>Pseudomonas</i> dosing)	30-50 <30	250-500 mg q12h 500 mg q24h
Clarithromycin (IR)	500 mg q12h	<30	250 mg q12h
*Entecavir	0.5-1 mg q24h	30-49 10-29 <10	0.5-1 mg q48h 0.5-1 mg q72h 0.5-1 mg q 7 days
Ethambutol	15-25 mg/kg q24h	10-50 <10	15-25 mg/kg q24-36h 15-25 mg/kg q48h
*Famciclovir	500 mg q8h (Herpes zoster dosing)	40-59 20-39 <20	500 mg q12h 500 mg q24h 250 mg q24h
Fluconazole	200-800 mg q24h	<50	Same LD → 100-400 mg q24h
*Flucytosine	25 mg/kg/dose q6h	21-40 10-20 <10	25 mg/kg/dose q12h 25 mg/kg/dose q24h 25 mg/kg/dose q48h
Lamivudine	100 mg q24h (HBV treatment; See page 22 for dosing considerations for HIV)	30-49 15-29	100 mg LD → 50 mg q24h 100 mg LD → 25 mg q24h
Levofloxacin	500 mg q24h (most indications) 750 mg q24h (anti- <i>Pseudomonas</i> coverage)	20-49 <20 20-49 <20	500 mg LD → 250 mg q24h 500 mg LD → 250 mg q48h 750 mg q48h 750 mg LD → 500 mg q48h
Nitrofurantoin	100 mg q12h	30-60 <30	Limit to short course (≤ 7 days) Use not recommended
Oseltamivir	See dosing on page 15		
Trimethoprim/sulfamethoxazole (TMP/SMZ)	1 DS tablet q12h (most indications) 1 DS tablet q24h or 3x/week (PCP ppx)	15-30 15-30	1 DS tablet → 1 SS tablet q12h 1 SS tablet q24h or 3x/week
Valacyclovir	1000 mg q12h (HSV) or 1000 mg q8h (herpes zoster)	30-49 10-29 <10	1000 mg q12h 1000 mg q24h 500 mg q24h
*Valganciclovir	900 mg q12h (induction dose) 900 mg q24h (maintenance/prevention dose)	40-59 25-39 10-24 40-59 25-39 10-24	450 mg q12h 450 mg q24h 450 mg q48h 450 mg q24h 450 mg q48h 450 mg twice weekly

\* = Restricted to Infectious Diseases/Antimicrobial Stewardship

BOLD = Nonformulary; requires PADR consult approval

LD = loading dose

# OVERVIEW OF COMMON ANTIRETROVIRAL THERAPY

22

When used for treatment or post-exposure prophylaxis (PEP), a complete antiretroviral regimen contains medications from 2, 3, or more drug classes to prevent the development of resistance.

- Many ARV tablets are combination pills which include more than 1 medicine in a single pill.
- In general, if 1 medication needs to be held, the entire regimen should be held. Infectious Diseases and Antimicrobial Stewardship teams can assist with choosing an alternative regimen.
- The preferred PEP regimen is raltegravir PLUS emtricitabine/tenofovir x28-30 days. PEP must be started within 72 hours of possible exposure.

When used for pre-exposure prophylaxis (PrEP) only two medications are needed.

- The only FDA-approved options for PrEP are emtricitabine/tenofovir alafenamide or emtricitabine/tenofovir disoproxil.

Any initiation of/change to antiretroviral therapy must be done in consultation with Infectious Diseases team. Use care to ensure the entire outpatient regimen is ordered upon admission/discharge from the hospital.

Refer to the "Crushing and Liquid ARV Formulations" link at [www.hivclinic.ca](http://www.hivclinic.ca) for an up-to-date list

Medication	Brand Name	Typical Dose	Renal Dose Adjustment per eGFR	Can be crushed, split, or opened?	Miscellaneous Info
Abacavir 300 mg TAB	Ziagen®	600 mg q24h	N/A	Yes	Separate from cations.
Abacavir/lamivudine/dolutegravir 600/300/50 mg TAB	Trumeq®	1 tablet q24h	<50: Not recommended, use individual components ≥50: Not recommended	Yes	Separate from cations.
Bictegravir/emtricitabine/tenofovir AF 50/200/25 mg TAB	Biktarvy®	1 tablet q24h	<30: Not recommended	No	Not recommended
Darunavir 600, 800 mg TAB	Prezista®	600 mg q12h or 800 mg q24h	N/A	No	Give with food. Give with ritonavir or co-cistat booster.
Darunavir/cobicistat 800/150 mg TAB	Prizcoxin®	1 tablet q24h	N/A	No	Give with food.
Dolutegravir/avir 50 mg TAB	Tivicay®	1 tablet q24h	<30: Use with caution, consider alternative alternative	Yes	Separate from cations.
Dolutegravir/lamivudine 50/300 mg TAB	Dotarivo®	1 tablet q24h	<50: Not recommended alternative	No	Separate from cations.
Dolutegravir/rilpivirine 50/25 mg TAB	Juluca®	1 tablet q24h	N/A	No	Give with food. Separate from cations.
Doravirine 100 mg TAB	Pifetro®	1 tablet qHS	<50: Not recommended	No	Use in place of Atreipla®. Give on empty stomach at bedtime.
Efavirenz 600 mg TAB	Sustiva®	1 tablet qHS	N/A	No	Give with food. Separate from cations.
Efavirenz/lamivudine/tenofovir DF 600/300/300 mg TAB	Genvoya®	1 tablet q24h	<30: Not recommended	No	Give with food. Separate from cations.
Emtricitabine/cobicistat/emtricitabine/tenofovir AF 150/150/200/10 mg TAB	Stribild®	1 tablet q24h	<70: Initial use not recommended <50: Cont'd use not recommended	No	Give with food. Separate from cations.
Emtricitabine/tenofovir AF 200/25 mg TAB	Descovy®	1 tablet q24h	<30: Not recommended	No	Avoid use for PrEP for patients engaging in receptive vaginal intercourse.
Emtricitabine/tenofovir DF 200/300 mg TAB	Truvada®	1 tablet q24h	<60: Not recommended (PrEP only) <30: 1 tablet q48h	Yes	Used for pre- or post-exposure prophylaxis (PrEP/PEP).
Emtricitabine/rilpivirine/tenofovir AF 200/25/25 mg TAB	Odiefsey®	1 tablet q24h	<30: Not recommended	No	Give with food. Do not use if HIV VL>100K or CD4<200
Emtricitabine/tenofovir DF 200/300 mg TAB	Intelence®	200 mg q12h	N/A	Yes	Give AFTER meals
Etravirine 200 mg TAB	Epivir®	300 mg q24h	30-49: 150 mg q24h 5-14: 150 mg x1 → 50 mg q24h <5: 50 mg x1 → 25 mg q24h	Yes	
Lamivudine 100, 150 mg TAB	Kaletra®	2 tablets q12h	N/A	No	400 and 600 mg tablets should not be interchanged
Lopinavir/ritonavir 200/50 mg TAB	Isentress®	400 mg q12h 1200 mg q24h	-	No	Tablets and capsules NOT bioequivalent. Used to "boost" protease inhibitors.
Raltegravir 400 mg TAB	Raltegravir 500 mg TAB	-	-	-	Give with food
Ritonavir 100 mg TAB	Norvir®	100 to 400 mg per day in 1-2 divided doses	N/A	No	Give with food
Tenofovir AF 25 mg TAB	Vemlidy®	25mg q24h	<15: Not recommended	No	
Tenofovir DF 300 mg TAB	Viread®	30-49: 300 mg q48h 10-29: 300 mg q24h	30-49: 300 mg q48h 10-29: 300 mg q24h	Yes	

# ALTERNATIVE DOSING: SELECT ANTIMICROBIALS

23

## Cefepime (see SOP 119-42)

Indication	<i>CrCl &gt;50 ml/min</i>	<i>CrCl 30-50 ml/min</i>	<i>CrCl 10-29 ml/min</i>	<i>CrCl &lt;10 or HD</i>
CNS infections Neutropenic fever Elevated cefepime MIC (4-8)* Empiric HAP/VAP coverage	2 g q8h	2 g q12h	1 g q12h	1 g q24h
CAP (not <i>Pseudomonas</i> ) Mild to moderate UTI	1 g q12h	1 g q24h	1 g q24h	500 mg q24h
All other indications	1 g q6h	1 g q8h	1 g q12h	1 g q24h

\*For *Enterobacteriaceae*

## Meropenem (see SOP 119-42)

Indication	<i>CrCl &gt;50 ml/min</i>	<i>CrCl 26-50 ml/min</i>	<i>CrCl 10-25 ml/min</i>	<i>CrCl &lt;10 or HD</i>
CNS infections Neutropenic fever Elevated carbapenem MIC (>4)	2 g q8h	2 g q12h	1 g q12h	1 g q24h
Uncomplicated UTI	500 mg q8h	500 mg q12h	250 mg q12h	500 mg q24h
All other indications	500 mg q6h	500 mg q8h	500 mg q12h	500 mg q24h

## Piperacillin/tazobactam (see SOP 119-24)

<i>CrCl</i>	<i>Indication</i>	<i>Intermittent Dosing</i>	<i>Extended Infusion Dosing</i>
>20 ml/min and piperacillin/tazobactam intermediate or resistant <i>Pseudomonas aeruginosa</i> (breakpoints: S<16, I 32-64, R >128)		Not recommended	4.5 g IV q6h over 3 hours (recommend additional anti- <i>Pseudomonal</i> coverage)
>40 ml/min	HAP or critical illness	4.5 g IV q6h over 60 min	4.5 g IV q8h over 4 hours
	Other indications	3.375 g IV q6h over 30 min	3.375 g IV q8h over 4 hours
20 – 40 ml/min	HAP or critical illness	3.375 g IV q6h over 30 min	3.375 g IV q8h over 4 hours
	Other indications	2.25 g IV q6h over 30 min	2.25 g IV q8h over 4 hours
<20 ml/min	HAP or critical illness	2.25 g IV q6h over 30 min	2.25 g IV q8h over 4 hours
	Other indications	2.25 g IV q8h over 30 min	2.25 g IV q12h over 4 hours
Hemodialysis (HD) and peritoneal dialysis (PD)	HAP or critical illness	2.25 g IV q8h over 30 min & 0.75 g post-dialysis	2.25 g IV q8h over 4 hours
	Other indications	2.25 g IV q12h over 30 min & 0.75 g post-dialysis	2.25 g IV q12h over 4 hours
Continuous Renal Replacement Therapy (CRRT)	Follow CrCl > 40 ml/min dosing	Follow CrCl > 40 ml/min dosing	

# GUIDE TO HOME IV THERAPY

24

## Home IV Antimicrobial Procedures

- Physician enters the "HOME IV ANTIBIOTIC NOTE" by searching by progress note title and then fills out the required template. Choose "CVS/Coram" under the Infusion Company drop down list
- Any changes to the regimen must be documented on the above note and cosigned by the attending physician
- Orders must be entered and signed by 12pm if patient is leaving the same day
- Transition patients to ideal dosing schedule → 08:00 for Q24H dosing, 08:00/20:00 for Q12H dosing
- Schedule labs and serum monitoring for early in the week (Mondays through Wednesdays)

## CVS/Coram Specialty Infusion Contact Information

- Phone (813) 639-4500; fax (877) 602-6777
- Case managers: Katie Sullivan & Jessica Yanta

## Types of Home Infusion:

IV Push	Continuous Infusion Pump												
<ul style="list-style-type: none"><li>Convenient</li><li>Does NOT restrict mobility</li><li>Medication is provided in a pre-filled syringe</li><li>The patient or caregiver manually infuses the medication over several minutes</li></ul> <p>Antimicrobials available via this method:</p> <table><tbody><tr><td>• Aztreonam</td><td>• Cefoxitin</td></tr><tr><td>• Cefazolin</td><td>• Ceftazidime</td></tr><tr><td>• Cefepime</td><td>• Ceftriaxone</td></tr><tr><td>• Cefotetan</td><td>• Daptomycin</td></tr></tbody></table>	• Aztreonam	• Cefoxitin	• Cefazolin	• Ceftazidime	• Cefepime	• Ceftriaxone	• Cefotetan	• Daptomycin	<ul style="list-style-type: none"><li>Convenient</li><li>Does NOT restrict mobility</li><li>Medication is provided in either a pre-filled syringe or an IV bag</li><li>The patient or caregiver places the medication in the pump which then infuses over a programmed amount of time, usually 24 hours</li><li>The medication/pump may be placed in a backpack or fanny pack so the patient can perform daily activities</li></ul> <p>Antimicrobials available via this method:</p> <table><tbody><tr><td>• Cefazolin</td><td>• Penicillin G</td></tr><tr><td>• Oxacillin</td><td>• Piperacillin/tazobactam</td></tr></tbody></table>	• Cefazolin	• Penicillin G	• Oxacillin	• Piperacillin/tazobactam
• Aztreonam	• Cefoxitin												
• Cefazolin	• Ceftazidime												
• Cefepime	• Ceftriaxone												
• Cefotetan	• Daptomycin												
• Cefazolin	• Penicillin G												
• Oxacillin	• Piperacillin/tazobactam												

Elastomeric Pump	Gravity IV																																
<ul style="list-style-type: none"><li>Convenient</li><li>Does NOT restrict mobility</li><li>Medication provided in a pre-filled eclipse ball</li><li>The patient or caregiver connects the pump which infuses the medication over a programmed amount of time</li><li>The ball may be placed in a pocket or backpack so the patient can perform daily activities</li></ul> <p>Antimicrobials available via this method:</p> <table><tbody><tr><td>• Amikacin</td><td>• Daptomycin</td></tr><tr><td>• Aztreonam</td><td>• Ertapenem</td></tr><tr><td>• Cefazolin</td><td>• Gentamicin</td></tr><tr><td>• Cefepime</td><td>• Oxacillin</td></tr><tr><td>• Cefotetan</td><td>• Penicillin G</td></tr><tr><td>• Cefoxitin</td><td>• Piperacillin/tazobactam</td></tr><tr><td>• Ceftazidime</td><td>• Tobramycin</td></tr><tr><td>• Ceftriaxone</td><td>• Vancomycin</td></tr><tr><td>• Clindamycin</td><td></td></tr></tbody></table>	• Amikacin	• Daptomycin	• Aztreonam	• Ertapenem	• Cefazolin	• Gentamicin	• Cefepime	• Oxacillin	• Cefotetan	• Penicillin G	• Cefoxitin	• Piperacillin/tazobactam	• Ceftazidime	• Tobramycin	• Ceftriaxone	• Vancomycin	• Clindamycin		<ul style="list-style-type: none"><li>NOT convenient</li><li>Restricts mobility</li><li>Medication is provided in an IV bag</li><li>Patient or caregiver must spike the bag, prime the line, set rate, and hang the medication on an IV pole for the duration of infusion. In some cases, the patient or caregiver must also reconstitute the medication</li></ul> <p>Antimicrobials available via this method:</p> <table><tbody><tr><td>• Ampicillin</td><td>• Levofloxacin</td></tr><tr><td>• Ampicillin/sulbactam</td><td>• Linezolid</td></tr><tr><td>• Azithromycin</td><td>• Meropenem</td></tr><tr><td>• Ceftaroline</td><td>• Metronidazole</td></tr><tr><td>• Ciprofloxacin</td><td>• Minocycline</td></tr><tr><td>• Doxycycline</td><td>• Tigecycline</td></tr><tr><td>• Imipenem/cilastatin</td><td>• TMP/SMZ</td></tr></tbody></table>	• Ampicillin	• Levofloxacin	• Ampicillin/sulbactam	• Linezolid	• Azithromycin	• Meropenem	• Ceftaroline	• Metronidazole	• Ciprofloxacin	• Minocycline	• Doxycycline	• Tigecycline	• Imipenem/cilastatin	• TMP/SMZ
• Amikacin	• Daptomycin																																
• Aztreonam	• Ertapenem																																
• Cefazolin	• Gentamicin																																
• Cefepime	• Oxacillin																																
• Cefotetan	• Penicillin G																																
• Cefoxitin	• Piperacillin/tazobactam																																
• Ceftazidime	• Tobramycin																																
• Ceftriaxone	• Vancomycin																																
• Clindamycin																																	
• Ampicillin	• Levofloxacin																																
• Ampicillin/sulbactam	• Linezolid																																
• Azithromycin	• Meropenem																																
• Ceftaroline	• Metronidazole																																
• Ciprofloxacin	• Minocycline																																
• Doxycycline	• Tigecycline																																
• Imipenem/cilastatin	• TMP/SMZ																																

# VANCOMYCIN DOSING & MONITORING

25

Remember to place a Therapeutic Drug Monitoring (TDM) consult for patients on vancomycin or aminoglycosides. Pharmacy service will dose, monitor, and adjust the regimen as needed. Be sure to include your indication for use!

## **Empiric Dosing:**

### **1. Determine pharmacokinetic targets based on indication.**

Goal AUC (mg*hr/L)	Infection Classification	Example(s)
400 – 450	Mild	Uncomplicated cystitis, cellulitis without systemic infection
450 – 500	Moderate	Pyelonephritis, cellulitis with systemic infection
500 – 550	Severe	Pneumonia, bacteremia, endocarditis, sepsis, osteomyelitis

### **2. Input patient data into the vancomycin calculator.**

- (S:) Pharmacy → Clinical → Vancomycin Kinetics → Vancomycin Calculator
- Select the volume of distribution (Vd) factor and clearance method suggested by the calculator
- If unable to access the above calculator, use the calculator here: <https://www.vancopk.com/>

### **3. Select a one-time loading dose (if necessary)**

- 20-35 mg/kg actual body weight if BMI <30 kg/m<sup>2</sup> or 20-25 mg/kg actual body weight if BMI ≥30 kg/m<sup>2</sup>
- Consider a loading dose for patients with severe infection
- Maximum single loading dose = 3000 mg

### **4. Select maintenance dose and dosing interval**

- Choose the 'best fit' or 'next best fit' regimen as suggested by the calculator
- Maximum single maintenance dose = 2000 mg.
- Minimum infusion rate is 30 minutes per each 500 mg. Consider extending infusion time in the setting of Redman's syndrome.

## **Monitoring:**

### **Determine if using traditional trough-based or AUC-based monitoring**

Monitoring is recommended for all patients with serious MRSA infections, those at high risk for nephrotoxicity, patients with variable renal function, or those receiving more than 3-5 days of therapy. Monitoring is typically not necessary for patients receiving vancomycin for surgical prophylaxis, uncomplicated UTI, cellulitis without bacteremia, or other indications where anticipated duration of therapy is less than 3-5 days.

- Trough-based monitoring
- Meningitis or CNS infection
  - Variable renal function, HD, or CRRT

- AUC-based monitoring
- All other indications

Trough level only

Both peak and trough levels

What levels should be drawn?

Obtain level at steady state\*

- Draw trough 30 minutes prior to next dose

\*For HD patients, obtain a pre-HD level prior to 2<sup>nd</sup> or 3<sup>rd</sup> supplemental dose. Consider obtaining with morning labs if patient has no residual renal function.

Obtain levels at steady state<sup>§</sup>

- Draw peak 2 hours after the infusions ENDs
- Draw trough 30 minutes prior to next dose

<sup>§</sup>May draw levels after the 1<sup>st</sup> maintenance dose if a loading dose of ≥20 mg/kg actual body weight was given.

Once at goal, surveillance troughs should be monitored weekly (unless pharmacokinetic parameters change)

What are the target/goals?

- Trough 10-15 mcg/mL: UTIs and SSTIs
- Trough 15-20 mcg/mL: All other indications

- AUC 400-550 mg\*hr/L and
- Trough 10-20 mcg/mL

# AMINOGLYCOSIDE DOSING & MONITORING: CONVENTIONAL

26

## Conventional Dosing

1. Calculate the patient's ideal body weight (IBW). Use equation on page #18.
2. Calculate an initial dose. Use IBW to calculate the dose unless...
  - Patient weighs less than their IBW, then use actual body weight (ABW)
  - Patient weighs more than 120% of their IBW, then use adjusted body weight (AdBW)

Tobramycin and Gentamicin	1 - 2.5 mg/kg
Amikacin	5 - 7.5 mg/kg

3. Calculate the patient's creatinine clearance. Use equation on page #18. Consider using cystatin-C to estimate creatinine clearance in spinal cord injury patients (<http://www.touchcalc.com/calculators/cystatin>)
4. Select a dosing interval based on the creatinine clearance.

Creatinine clearance (mL/min)	Suggested Dosing Interval
>60	Q8h
40-59	Q12h
20-39	Q24h
<20	Loading dose then follow levels
Dialysis	See dialysis dosing on pages 20

5. Once at steady state, draw a trough level 30 minutes prior to the next infusion and a peak level 30 minutes after the infusion has ended. Adjust the regimen as necessary.

Infection	Desired Peak (mcg/mL)		Desired Trough (mcg/mL)	
	Tobra/gent	Amikacin	Tobra/gent	Amikacin
Pneumonia	8-10	25-35	<1	<4-8
Other gram negative infections	6-8	25-35	<1	<4-8
Cystitis, synergy in gram positive infections	3-5	20-25	<1	<4-8

## Useful PK Equations (when levels available)

Patient-specific Ke:

$$Ke = \frac{\ln(C_{\text{peak measured}} / C_{\text{trough measured}})}{\Delta t}$$

$\Delta t$  = time between measured peak and measured trough OR time interval between 2 levels with no doses given in between

Half-life:

$$t_{1/2} = 0.693 / Ke$$

True peak/trough:

$$\text{True } C_{\text{peak}} = \frac{C_{\text{peak measured}}}{e^{-(Ke)(t)}}$$

Where  $t$  = the time peak was drawn minus time infusion ended

$$\text{True } C_{\text{trough}} = [C_{\text{trough measured}}] \times [e^{-(Ke)(t)}]$$

Where  $t$  = time infusion began minus time trough was drawn

Volume of distribution:

$$Vd = (\text{dose}/t/Ke) \times \frac{1 - e^{-(Ke)(t)}}{C_{\text{true peak}} - (C_{\text{true trough}} \times e^{-(Ke)(t)})}$$

Where  $t$  = the time of infusion (usually 0.5 or 1 hour)

New dosing interval

$$\tau = \frac{-1}{Ke} \times [\ln(C_{\text{trough desired}} / C_{\text{peak desired}}) + t]$$

New dose

$$\text{New dose} = (Ke \times Vd \times C_{\text{peak desired}}) \times \frac{1 - e^{-(Ke)(t)}}{1 - e^{-(Ke)(\tau)}}$$

Where  $t$  = the time of infusion (usually 0.5 or 1 hour)

# AMINOGLYCOSIDE DOSING & MONITORING: EXTENDED INTERVAL

27-28

## Determine if using Barnes-Jewish or Hartford Hospital nomogram

Extended interval dosing is not recommended for burn, pregnant, or trauma patients nor those with endocarditis, ascites, or unstable/fluctuating renal function. Avoid extended interval dosing when using aminoglycosides for synergy and for patients with ESRD/on dialysis

### Barnes-Jewish Nomogram

- Use for most patients
- Gentamicin/tobramycin 5 mg/kg
- Amikacin 15 mg/kg

### Hartford Hospital nomogram

- Use for pneumonia or severe sepsis
- Gentamicin/tobramycin 7 mg/kg

## Calculate an initial dose

See ideal body weight (IBW) and adjusted body weight (AdBW) equations on page 18

### Use ideal body weight, unless...

- Patient weighs less than IBW, then use actual body weight
- Patient weighs >20% IBW, then use adjusted body weight

### Use actual body weight, unless...

- Patient weighs >20% IBW, then use adjusted body weight

## Draw a post-dose level

Draw a level 6-14 hours AFTER the infusion has ended

Draw a level 6-14 hours after the START of the infusion

## Plot the level on the appropriate nomogram and adjust patient's dosing interval

Barnes-Jewish nomograms to the right

Hartford Hospital nomogram below

14

13

12

11

10

9

8

7

6

5

4

3

2

Gentamicin/Tobramycin 7 mg/kg

Q48h

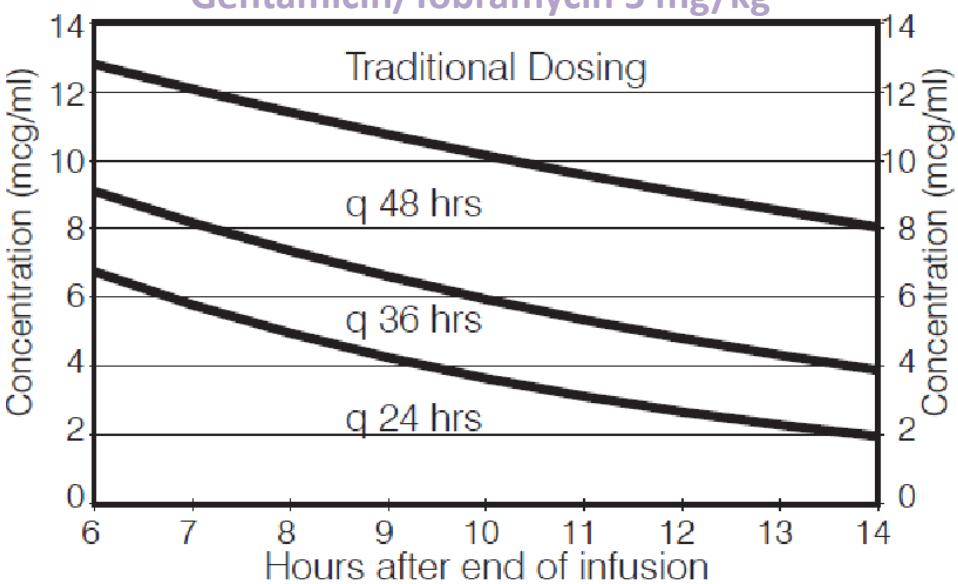
Q36h

Q24h

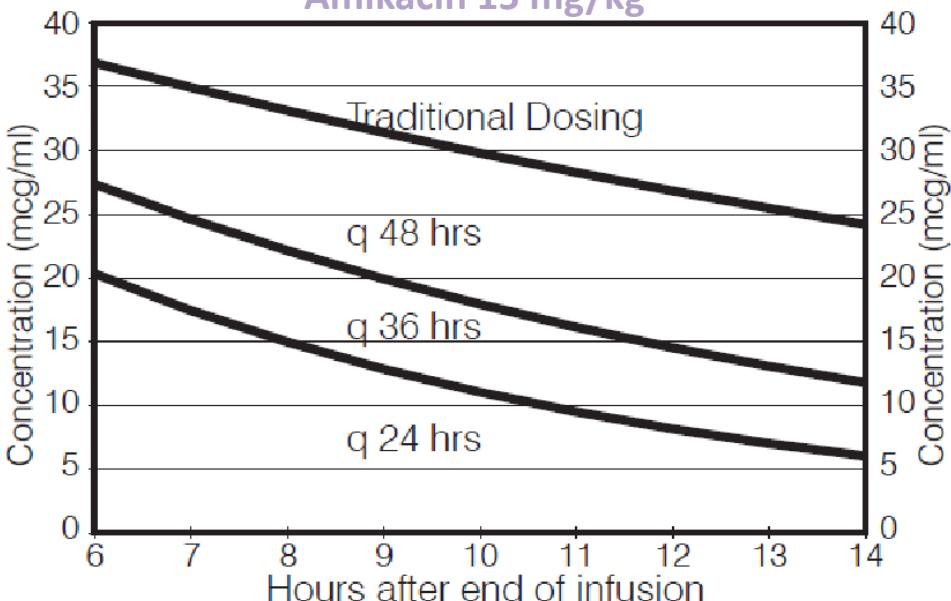
6 7 8 9 10 11 12 13 14

Time between start of infusion and sample draw (hrs)

### Gentamicin/Tobramycin 5 mg/kg



### Amikacin 15 mg/kg



Bailey TC, et al. A Meta-Analysis of Extended-Interval Dosing Versus Multiple Daily Dosing of Aminoglycosides. *Clin Infect Dis*. 1997;24:786-95

Nicolau DP, et al. Experience with a Once-Daily Aminoglycoside Program Administered to 2,184 Adult Patients. *Antimicrob Agents Chemother*. 1995;39(3):650-655

## GRAM POSITIVE ISOLATES (2018)

**Gram Positive  
Organisms 2018**

Gram Positive Organisms 2018		Sensitivity Data (%)					
Antibiotic	Susceptible	Intermediate	Resistant	ESBL	MIC <sub>50</sub>	MIC <sub>90</sub>	Comments
AMPICILLIN	99	23					LUGDUNENSIS
CIPROFLOXACIN	73	9	50	71	47	50	STAPHYLOCOCCUS
CLINDAMYCIN			70	71	64	52	HOMINIS
DAPTOMYCIN	100		100	100	100	100	HAEMOLYTICUS
DOXYCYCLINE			96				STAPHYLOCOCCUS
ERYTHROMYCIN	10	2	34	68	35	37	EPIDERMIDS
GENTAMICIN			99	82	92	92	STAPHYLOCOCCUS
LEVOFLOXACIN	75	11	51	71	48	55	AUREUS
LINEZOLID	92	96	100	100	100	100	CAPITUS
NITROFURANTOIN			69	93	77	79	STAPHYLOCOCCUS
OXACILLIN			46	71	43	55	HOMINIS
RIFAMPIN			99	100	98	95	HAEMOLYTICUS
TETRACYCLINE	19	15	91	86	80	71	EPIDERMIDS
TIGECYCLINE	100	100	100	100	100	100	TETRACYCLINE
TRIMETHOPRIM/SULFA			87	93	57	82	TRIMETHOPRIM/SULFA
VANCOMYCIN	94	38	99	100	99	100	VANCOMYCIN

# GRAM NEGATIVE ISOLATES (2018)

30

## Gram Negative Organisms 2018

	AMIKACIN	AMPICILLIN	AMPICILLIN/SULBACTAM	CETEFAZOLIN	CEFERIME	CEFTAZIDIME	CEFTRIAXONE	CIPROFLOXACIN	ERTAPENEM	GENTAMICIN	IMIPENEM	LEVOFLOXACIN	NITROFURANTOIN	PIPERACILLIN/TAZOBACTAM	TOBRAMYCIN	TRIMETHOPRIM/SULFA
ACINETOBACTER BAUMLANNI COMPLEX	100	100	100	100	95	97	98	94	96	98	97	98	96	98	97	98
CITROBACTER FREUNDII	89	98	97	95	100	92	90	96	90	96	97	91	91	100	90	95
CITROBACTER AEROGENES	69	98	98	82	88	82	88	99	88	99	95	91	97	91	90	92
ENTEROBACTER CLOACAE	81	89	82	82	88	82	88	99	88	82	88	82	88	87	87	88
ESCHERICHIA COLI	63	98	98	88	82	88	88	99	88	98	95	91	97	91	95	95
KLEBSIELLA OXYTOSCA	13	81	86	88	82	88	82	99	88	99	84	95	91	97	100	92
KLEBSIELLA PNEUMONIAE	56	95	98	95	94	66	97	85	67	95	65	52	81	81	97	97
MORGANELLA MORGANI	95	98	98	95	94	96	97	95	97	98	99	98	99	99	99	99
PROVIDENCIA STUARTII	95	98	98	95	94	97	98	95	97	98	99	98	99	99	99	99
PSEUDOMONAS AERUGINOSA	95	98	98	95	94	97	98	95	97	98	99	98	99	99	99	99
PROTEUS MIRABILIS	95	98	98	95	94	97	98	95	97	98	99	98	99	99	99	99
PROVIDENCIA STUARTII	95	98	98	95	94	97	98	95	97	98	99	98	99	99	99	99
SALMOSELLA MARCESCENS	95	98	98	95	94	97	98	95	97	98	99	98	99	99	99	99
STENOTROPHOMONAS MALTOPHILA	95	98	98	95	94	97	98	95	97	98	99	98	99	99	99	99

At the time of printing, an updated antibiogram with 2019 isolates was not yet available. Please refer to the antibiogram on the Intranet homepage for most updated version.

Intranet → Clinical Links (on right side menu) → Hospital Antibiogram

# IV TO PO CONVERSION

31

## Candidates for Antimicrobial Step-Down Therapy

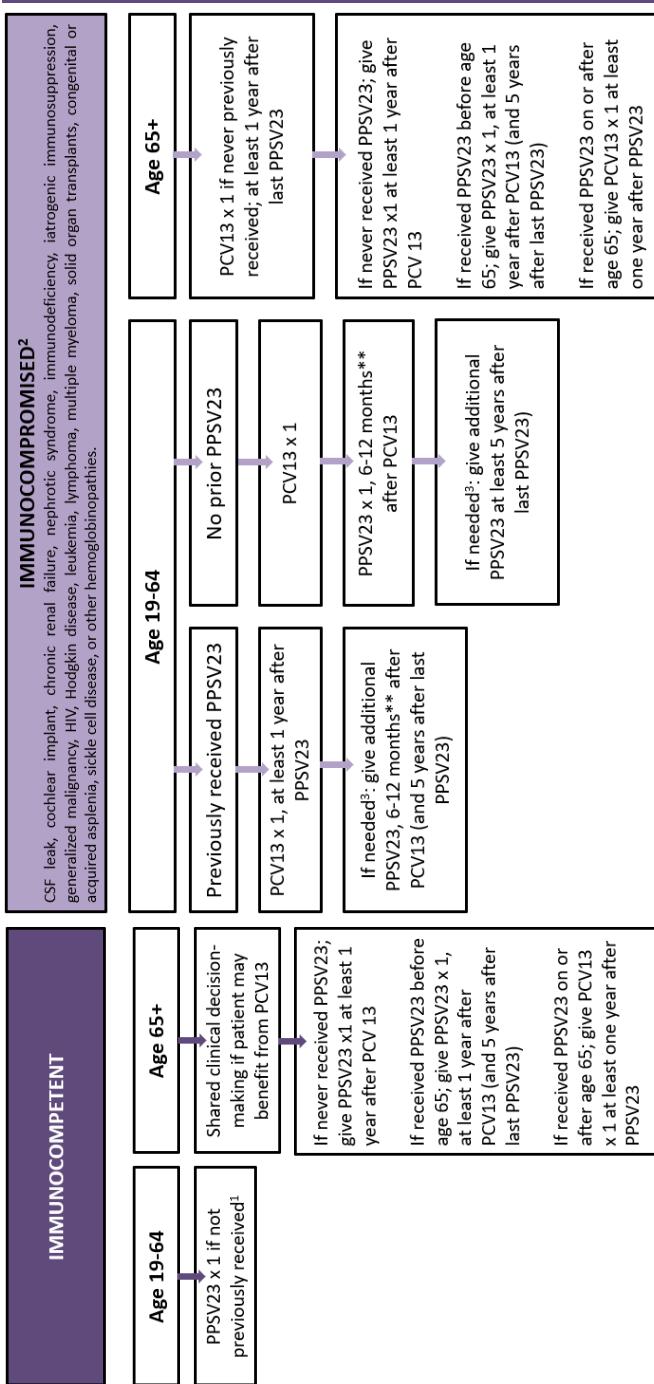
- The infection is treatable with oral antimicrobial therapy and the indications and spectrum of activity are identical/similar between drugs
- Patient is able to take medications by mouth or has a functional feeding tube in place
- Pt has been on IV therapy ≥48 hours and is showing overall clinical improvement
  - No fever
  - WBC within normal limits or trending down to normal range
  - Hemodynamically stable as evidenced by:
    - Temp. ≤38°C (100.4°F) and ≥36°C (96.8°F)
    - HR <90 beats/minute
    - RR <20 breaths/minute
    - Stable BP

## Contraindications for Antimicrobial Step-Down Therapy

- Active NPO order or inadequately absorbing PO medications
  - Vomiting
  - Malabsorption disorder
  - Mechanical swallowing disorder
  - On gastric suction or gastric paralytics
  - GI obstruction or abnormal motility/ileus
- Serious infections requiring prolonged IV antibiotic treatment including, but not limited to:
  - Meningitis
  - Osteomyelitis
  - Endocarditis
  - Bacteremia/sepsis
  - Liver abscess

IV	PO
Acyclovir 5 mg/kg q8h	Valacyclovir 1000 mg q12h
Acyclovir 10mg/kg q8h	Valacyclovir 1000 mg q8h
Ampicillin 1 g q6h	Amoxicillin 500 mg q8h
Ampicillin/sulbactam 1.5–3gm q8h	Amoxicillin/clavulanate 875/125 mg q12h (except <i>Acinetobacter</i> spp.)
Azithromycin 500 mg q24h	Same dose and interval
Cefazolin 1 g q8h	Cephalexin 500 mg q6h
Ceftriaxone 1-2g q24h	Cefdinir 300mg q12h (except <i>Morganella</i> spp.)
Ciprofloxacin 200 mg q12h	Ciprofloxacin 250 mg q12h
Ciprofloxacin 400 mg q12h	Ciprofloxacin 500 mg q12h
Ciprofloxacin 400 mg q8h	Ciprofloxacin 750 mg q12h
Clindamycin 600 mg q8h	Clindamycin 300 mg q8h
Clindamycin 900 mg q8h	Clindamycin 450 mg q8h
Doxycycline 100 mg q12h	Same dose and interval
Fluconazole 100–400 mg q24h	Same dose and interval
Levofloxacin 500–750 mg q24h	Same dose and interval
Linezolid 600 mg q12h	Same dose and interval
Metronidazole 500 mg q8h	Same dose and interval
Minocycline 100mg q12h	Same dose and interval
Rifampin 300 mg q12h	Same dose and interval

# PNEUMOCOCCAL VACCINE ALGORITHM



PPSV23 = 23-valent pneumococcal polysaccharide vaccine, Pneumovax®, Merck & Co., Inc.  
 PCV13 = 13-valent pneumococcal conjugate vaccine, Prevnar®, Pfizer Inc.

\*\* Ideally PCV13 and PCV13 should be given 6-12 months apart. However, minimum acceptable interval is 8 weeks

<sup>1</sup> Routine use of PPSV23 is not recommended for American Indian/Alaska Natives or other persons younger than age 65 unless they have underlying medical conditions that are PPSV23 indications

<sup>2</sup> PCV13 x 1 Exemption: Hematopoietic stem cell transplant (HSCT) patients should receive PCV13 x 3 doses before receiving PPSV23 (initial PCV13 dose 3-6 months after transplant, subsequent PCV13 doses every 2 months x 2 doses)

<sup>3</sup> Additional PPSV23 administrations: give a TOTAL of two PPSV23 doses to immunocompromised patients, EXCEPT for those with CSF leaks & cochlear implants (whom only receive 1 dose)

# PENICILLIN ALLERGIES

33

10% of the population reports a penicillin allergy but <1% of the whole population is truly allergic.



## Did You Know?

- Approximately 80% of patients with IgE-mediated penicillin allergy lose their sensitivity after 10 years.

### James A. Haley Veterans' Hospital Allergy Assessment Questionnaire

- 1 What is the name of the specific medication that you had a reaction to?
- 2 Where did you have the medication filled? If not at JAHVH, what type of medication did you have a reaction to (i.e. oral, IV, topical, etc.)?
- 3 How long ago did the reaction occur? How soon after taking the medication did you experience the reaction?
- 4 Describe the reaction that you experienced?
- 5 Did you require medical treatment for the reaction you experienced (i.e. did you come to the hospital, what medications did you receive)?
- 6 Were you taking any other new medications at the time of the reaction? If yes, are you still tolerating those medications?
- 7 Since having your initial reaction have you tolerated the same medication again or a medication in the same class? [provide medication examples]

## Can My Patient Receive a Penicillin or Cephalosporin?

Does patient report history of IgE mediated/Type I allergic reaction?

Characteristics of an IgE mediated/Type I reaction include:

- Reactions that occur immediately or usually within one hour
- Hives defined as multiple pink/red raised areas of skin that are intensely itchy
- Angioedema defined as localized edema without hives affecting the abdomen, face, extremities, genitalia, oropharynx, or larynx
- Wheezing and shortness of breath
- Anaphylaxis



- Avoid penicillins and cephalosporins.
- Consider penicillin skin testing (consult Allergy).
- De-sensitization required if no alternatives.



Yes or unknown

Was reported allergy severe?



Has patient tolerated any penicillins or cephalosporins since the allergy was reported?



- Consider use with same agent or agent with similar side chain (see table on next page)
- Consider penicillin skin testing (consult Allergy)



- Consider trial of beta-lactam agent with a side chain **dissimilar** to that of the offending agent with close monitoring
- Consider penicillin skin testing (consult Allergy)

# BETA LACTAM CROSS REACTIVITY

34

Mono	5 <sup>th</sup>	4 <sup>th</sup>	3 <sup>rd</sup>	2 <sup>nd</sup>	1 <sup>st</sup>	Penicillins	Cefazolin	Cefadroxil	Cephalexin	Cefuroxime	Cefotaxime	Ceftriaxone	Cephalosporine	Cefixime	Cefdinir	Cefuroxime	Cefotetan	Cefdinil	Ceftriaxone	Cefoperazone	Cefiderocol	Aztreonam
						Nafcillin																
						Oxacillin																
						Dicloxacillin																
						Penicillin G/VK																
						Piperacillin																
						Ampicillin																
						Amoxicillin																
						Oxacilline																
						Dicloxacilline																
						Nafcillin																

Adapted from Zagursky RJ et al. Cross-reactivity in  $\beta$ -lactam allergy. *J Allergy Clin Immunol Pract.* 2018;6:72-81

Blue shaded: Identical R<sub>1</sub> or R<sub>2</sub> structures  
 Purple shaded: Similar R<sub>1</sub> or R<sub>2</sub> structures or components (ring or branch chain moiety)  
 Blank: No R<sub>1</sub> or R<sub>2</sub> structural similarities

# HOSPITAL PRECAUTIONS

35

## Illnesses Requiring Contact Precautions:

### Agents/Diseases

- Adenovirus
- Antibiotic-resistant organisms: MRSA, VRE, or key multi-drug resistant gram negatives<sup>§</sup>
- Clostridium difficile ("C. diff")
- Diarrhea, acute, unknown cause
- Diphtheria, cutaneous form (Note: pharyngeal diphtheria requires droplet)
- Ebola fever
- Hepatitis A
- Herpes simplex, disseminated mucocutaneous
- Herpes zoster (shingles) localized in immunocompetent patient
- Lassa fever
- **Infection Control flags gram negatives of epidemiological significance**
- Lice
- Marburg virus
- Respiratory Syncytial virus (RSV)
- Rotavirus
- Scabies
- Shingles
- Staphylococcal skin/wound/burn infection, major
- Streptococcal skin/wound/burn infection, major

### Clinical Syndromes

- Abscess, draining major
- Cellulitis, acute (uncontained)
- Conjunctivitis, viral
- Decubitus ulcer, major, infected
- Diarrhea, acute, unknown cause\*
- Enterocolitis, *Clostridium difficile*
- Hemorrhagic fevers
- Impetigo
- SARS (also Airborne precautions)
- Shingles, localized in immunocompetent pt
- Skin/wound or burn infections, with hard to control drainage
- Wound infection, major

\*Gastroenteritis (acute diarrhea) in incontinent patients caused by any of the following: *Campylobacter*, *Clostridium difficile*, *Cryptosporidium*, *Escherichia coli*, *Giardia*, Hepatitis A virus, Norovirus, Rotavirus, *Salmonella*, *Shigella*, *Vibrio cholerae* (cholera), *Vibrio parahaemolyticus*, *Yersinia enterocolitica* —OR— if the cause of acute diarrhea is unknown, but an infectious cause is suspected.



## Illnesses Requiring Airborne Precautions:

- Tuberculosis (*Mycobacterium tuberculosis* complex)
- Chickenpox (Varicella) [also CONTACT precautions]
- Measles (Rubeola)
- Shingles (Herpes zoster) in an immunocompromised patient [also use CONTACT precautions]
- Disseminated shingles (Herpes zoster) [also use CONTACT precautions]
- SARS - Severe Acute Respiratory Syndrome [also use CONTACT precautions]
- Smallpox



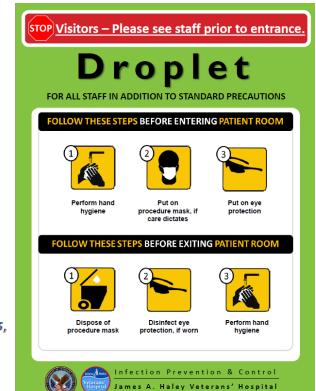
## Illnesses Requiring Droplet Precautions:

### Agents/Diseases

- *Corynebacteria diphtheriae*
- Diphtheria, pharyngeal
- Erythema infectiosum (Fifth Disease)
- Fifth Disease (parvovirus B19)
- *Haemophilus influenzae* meningitis
- *Haemophilus influenzae* pneumonia
- Influenza
- Mumps
- *Mycoplasma pneumoniae*
- *Neisseria meningitidis* bactemia
- *Neisseria meningitidis* meningitis
- *Neisseria meningitidis* pneumonia
- Parvovirus B19 (erythema infectiosum)
- Pertussis (whooping cough)
- Plague, pneumonic (*Yersinia pestis*)
- Rhinovirus (common cold)
- Rubella, (German measles)
- Viral Hemorrhagic Fevers (due to Lassa, Ebola, Marburg or Crimean-Congo fever viruses)
- Whooping cough (pertussis)
- *Yersinia pestis* pneumonia

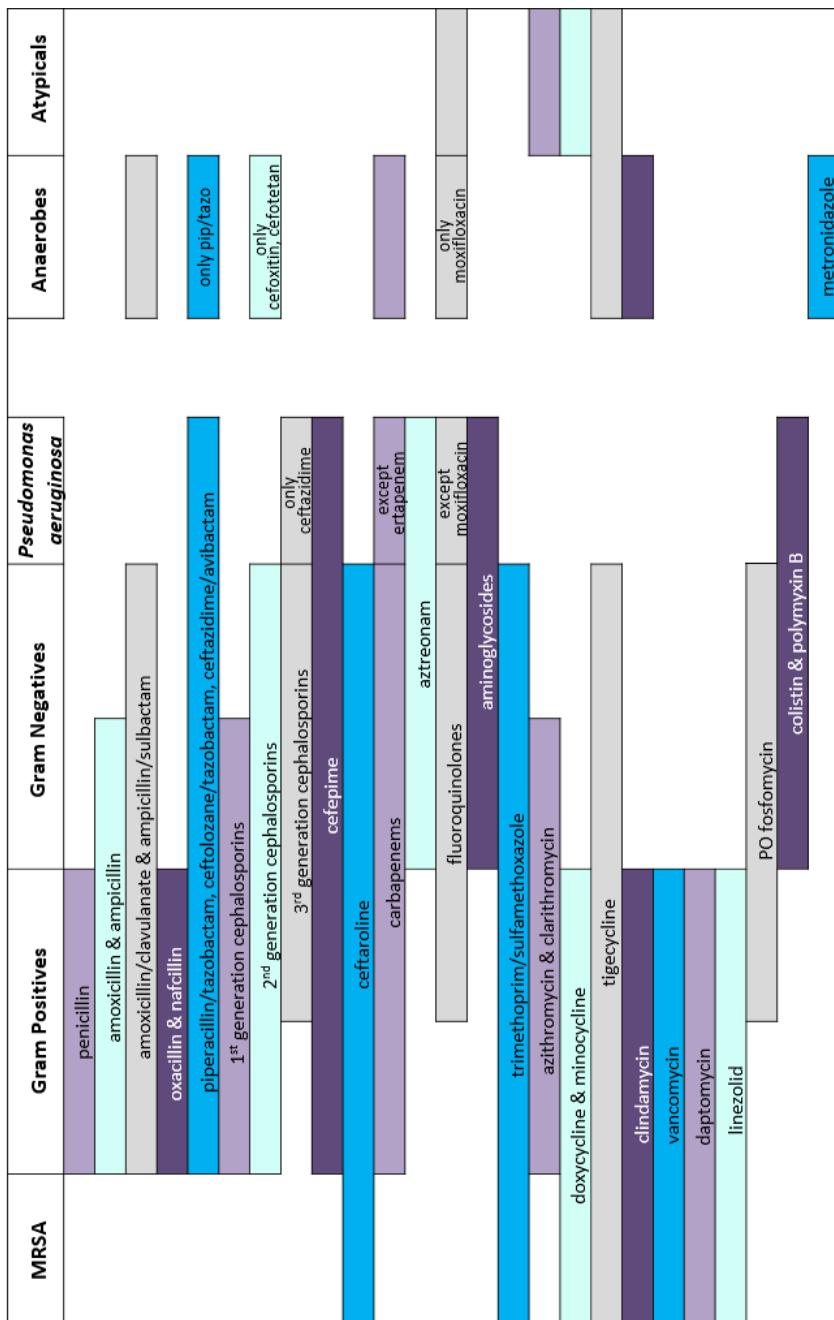
### Clinical Syndromes

- Epiglottitis, *Haemophilus influenzae*
- Influenza ("flu")
- Meningitis, *Neisseria meningitidis*
- Meningitis, *Haemophilus influenzae*
- Meningococcemia, *Neisseria meningitidis*
- Pneumonia – specifically with Adenovirus, *Mycoplasma pneumoniae*, *Neisseria meningitidis*, *Streptococcus Group A*
- Any "suspected" bacterial meningitis illness



# ANTIBIOTIC SPECTRUM OF ACTIVITY: AT A GLANCE

36



# SPECTRUM OF ACTIVITY

37-38

<b>Penicillins (penicillin V, penicillin G)</b>	Bactericidal
Good: <i>Treponema pallidum</i> , most <i>Streptococci</i>	
Moderate: <i>Enterococci</i>	
Poor: Everything else	
<b>Anti-staphylococcal Penicillins (nafcillin)</b>	Bactericidal
Good: <i>MSSA</i> , <i>Streptococci</i>	
Poor: Gram negative rods, <i>Enterococci</i> , anaerobes, <i>MRSA</i>	
<b>Aminopenicillins (amoxicillin, ampicillin)</b>	Bactericidal
Good: <i>Streptococci</i> , <i>Enterococci</i>	
Moderate: Enteric Gram-negative rods, <i>Haemophilus</i>	
Poor: <i>Staphylococci</i> , anaerobes, <i>Pseudomonas</i>	
<b>Beta-lactam/Beta-lactamase inhibitor combinations (piperacillin/tazobactam, amoxicillin/clavulanate, ampicillin/sulbactam)</b>	Bactericidal
Good: <i>MSSA</i> , <i>Streptococci</i> , <i>Enterococci</i> , anaerobes, Gram negative rods, <i>Pseudomonas aeruginosa</i> (only piperacillin/tazobactam)	
Moderate: Enteric Gram-negative rods with advanced beta-lactamases	
Poor: <i>MRSA</i> , enteric Gram-negative rods with extended-spectrum beta-lactamases (ESBL+)	
<b>Extended Spectrum Beta-lactam/Beta-lactamase inhibitor combinations (ceftazidime/avibactam, ceftolozane/tazobactam)</b>	Bactericidal
Good: <i>Pseudomonas aeruginosa</i> , enteric Gram-negative rods	
Moderate: Some <i>Streptococci</i>	
Poor: Anaerobes, <i>MRSA</i> , <i>MSSA</i> , <i>Acinetobacter</i>	
<b>1<sup>st</sup> generation cephalosporins (cefazolin, cephalexin)</b>	Bactericidal
Good: <i>MSSA</i> , <i>Streptococci</i>	
Moderate: Some enteric Gram-negative rods	
Poor: <i>Enterococci</i> , anaerobes, <i>MRSA</i> , <i>Pseudomonas</i>	
<b>2<sup>nd</sup> generation cephalosporins (cefuroxime, cefoxitin, cefotetan)</b>	Bactericidal
Good: Some enteric Gram negative rods, <i>Haemophilus</i> , <i>Neisseria</i>	
Moderate: <i>Streptococci</i> , <i>Staphylococci</i> , anaerobes (only cefoxitin and cefotetan)	
Poor: <i>Enterococci</i> , <i>MRSA</i> , <i>Pseudomonas</i>	
<b>3<sup>rd</sup> generation cephalosporins (ceftaxone, cefotaxime, ceftazidime, cefdinir, cefpodoxime, cefixime)</b>	Bactericidal
Good: <i>Streptococci</i> , <i>Pseudomonas</i> (ceftazidime only), enteric Gram negative rods	
Moderate: <i>MSSA</i> (except ceftazidime which is poor)	
Poor: <i>Enterococci</i> , <i>Pseudomonas</i> (except ceftazidime), <i>MRSA</i> , anaerobes	
<b>4<sup>th</sup> generation cephalosporins (cefepime)</b>	Bactericidal
Good: <i>MSSA</i> , <i>Streptococci</i> , <i>Pseudomonas</i> , enteric Gram negative rods	
Moderate: <i>Acinetobacter</i>	
Poor: <i>Enterococci</i> , anaerobes, <i>MRSA</i>	
<b>5<sup>th</sup> generation cephalosporins (ceftaroline)</b>	Bactericidal
Good: <i>MSSA</i> , <i>MRSA</i> , <i>Streptococci</i> , enteric Gram negative rods	
Moderate: <i>Acinetobacter</i> , <i>Enterococcus faecalis</i>	
Poor: <i>Pseudomonas</i> , <i>Enterococcus faecium</i> , anaerobes	
<b>Carbapenems (imipenem/cilastatin, meropenem, ertapenem)</b>	Bactericidal
Good: <i>MSSA</i> , <i>Streptococci</i> , anaerobes, Gram negative rods including ESBL+, <i>Pseudomonas/Acinetobacter</i> (not ertapenem)	
Moderate: <i>Enterococci</i> (not ertapenem)	
Poor: <i>MRSA</i>	
<b>Monobactams (aztreonam)</b>	Bactericidal
Good: <i>Pseudomonas</i> , most Gram negative rods	
Moderate: <i>Acinetobacter</i>	
Poor: Gram-positive organisms, anaerobes	
<b>Glycopeptides (vancomycin)</b>	Bactericidal*
Good: <i>MRSA</i> , <i>Streptococci</i> , <i>Clostridium difficile</i> (PO only), <i>MSSA</i>	
Moderate: <i>Enterococci</i> (*bacteriostatic)	
Poor: Gram-negative organisms	

<b>Fluoroquinolones (ciprofloxacin, levofloxacin, moxifloxacin)</b>		<b>Bactericidal</b>
Respiratory FQs – levofloxacin and moxifloxacin Good: Gram negative rods, <i>Haemophilus</i> , <i>Strep. pneumo</i> . Moderate: <i>Pseudomonas</i> (levofloxacin only), <i>MSSA</i> Poor: Anaerobes (except moxifloxacin which is moderate), <i>Enterococci</i>	Non-respiratory FQs – ciprofloxacin Good: Gram negative rods, <i>Haemophilus</i> Moderate: <i>Pseudomonas</i> , atypicals Poor: <i>Staphylococci</i> , <i>Strep pneumo</i> ., anaerobes, <i>Enterococci</i>	
<b>Aminoglycosides (gentamicin, tobramycin, amikacin)</b>		<b>Bactericidal</b>
Good: Gram-negatives Moderate: (in combination with a beta-lactam) <i>Staphylococci</i> , <i>Streptococci</i> , <i>Enterococci</i> Poor: Atypicals, anaerobes, gram-positive organisms		
<b>Tetracyclines (doxycycline, minocycline)</b>		<b>Bacteriostatic</b>
Good: Atypicals, <i>Rickettsia</i> , spirochetes Moderate: <i>Staphylococci</i> , <i>Strep pneumo</i> Poor: Gram negative rods, anaerobes, <i>Enterococci</i>		
<b>Glycylcycline (tigecycline)</b>		<b>Bacteriostatic</b>
Good: Atypicals, <i>Enterococci</i> , <i>Staphylococci</i> , <i>Strep pneumo</i> Moderate: Gram negative rods, anaerobes Poor: <i>Pseudomonas</i> , <i>Proteus</i> , <i>Providencia</i>		
<b>Macrolides (azithromycin, erythromycin, clarithromycin)</b>		<b>Bacteriostatic</b>
Good: Atypicals, <i>Haemophilus</i> , <i>M. catarrhalis</i> , <i>H. pylori</i> , <i>Mycobacteria</i> Moderate: <i>Strep pyogenes</i> Poor: <i>Staphylococci</i> , Gram negative rods, anaerobes, <i>Enterococci</i>		
<b>Oxazolidinone (linezolid)</b>		<b>Bacteriostatic</b>
Good: <i>MRSA</i> , <i>Streptococci</i> , <i>Enterococci</i> , <i>Nocardia</i> , <i>MSSA</i> Moderate: Some atypicals, some <i>Mycobacteria</i> Poor: Gram-negatives, anaerobes		
<b>Nitroimidazole (metronidazole)</b>		<b>Bactericidal</b>
Good: Anaerobes, protozoa Moderate: <i>H. pylori</i> Poor: Aerobes, anaerobes from human mouth flora		
<b>Nitrofurans (nitrofurantoin and fosfomycin)</b>		<b>Bactericidal</b>
Good: <i>E. coli</i> , <i>Staphylococcus saprophyticus</i> Moderate: <i>Citrobacter</i> , <i>Klebsiella</i> , <i>Proteus</i> , <i>Enterococci</i> , <i>Pseudomonas</i> (fosfomycin), <i>Serratia</i> (fosfomycin) Poor: <i>Acinetobacter</i>		
<b>Streptogramin (quinupristin/dalfopristin)</b>		<b>Bactericidal (<i>S. aureus</i>) Bacteriostatic (<i>E. faecium</i>)</b>
Good: <i>MSSA</i> , <i>MRSA</i> , <i>Streptococci</i> , <i>Enterococcus faecium</i> Poor: <i>Enterococcus faecalis</i> , Gram negative rods		
<b>Cyclic Lipopeptides (daptomycin)</b>		<b>Bactericidal</b>
Good: <i>MRSA</i> , <i>Streptococci</i> , <i>MSSA</i> , <i>Enterococci</i> Poor: Gram negative rods		
<b>Folate Antagonists (trimethoprim/sulfamethoxazole)</b>		<b>Bacteriostatic</b>
Good: <i>Staphylococcus aureus</i> , <i>Haemophilus</i> , <i>Stenotrophomonas</i> , <i>Listeria</i> , <i>Nocardia</i> Moderate: Gram negative rods, <i>Strep pneumo</i> , <i>Salmonella</i> , <i>Shigella</i> Poor: <i>Pseudomonas</i> , <i>Enterococci</i> , anaerobes		
<b>Lincosamides (clindamycin)</b>		<b>Bacteriostatic</b>
Good: Gram-positive anaerobes Moderate: <i>Staphylococcus aureus</i> , Gram-negative anaerobes, Poor: <i>Enterococci</i> , Gram-negative aerobes		
<b>Polymyxins (colistimethate sodium, polymyxin B)</b>		<b>Bactericidal</b>
Good: Multidrug resistant Gram negative rods Moderate: <i>Stenotrophomonas maltophilia</i> Poor: Gram-positive organisms, anaerobes		

# AVAILABLE ANTIMICROBIALS

39-40

<b>ANTIBIOTICS</b>	
Amikacin 250mg/mL INJ	Metronidazole 250 mg TAB
Amoxicillin 250, 500 mg CAP	Metronidazole 500 mg/100mL INJ
Amoxicillin 250 mg/5 mL ORAL SUSP	Minocycline 100 mg CAP
Amoxicillin/clavulanate 875/125mg TAB	*Minocycline 100 mg INJ
Amox/clav 250/62.5 mg/5 mL ORAL SUSP	Moxifloxacin 400 mg INJ/TAB
Ampicillin 1, 2 g INJ	Nafcillin 1, 2 g INJ
Ampicillin/sulbactam 1.5, 3 g INJ	Nitrofurantoin 100 mg CAP
Azithromycin 250, 500, 600 mg TAB	Penicillin G benzathine 1.2 million unit INJ
Azithromycin 500mg INJ	Penicillin G potassium 5 million unit INJ
Aztreonam 1, 2 g INJ	Penicillin VK 500 mg TAB
Cefadroxil 500 mg CAP	Piperacillin/tazo 2.25, 3.375, 4.5 g INJ
Cefazolin 500 mg, 1 g INJ	*Polymyxin B 500,000 IU INJ
Cefdinir 300 mg CAP	Tigecycline 50mg INJ
Cefepime 1, 2 g INJ	Tobramycin 80mg/2mL INJ
Cefotaxime 1, 2 g INJ	*Tobramycin 300 mg/5 mL INHALATION
Cefotetan 1, 2 g INJ	TMP/SMZ 800/160 mg TAB/INJ
Cefoxitin 1, 2 g INJ	TMP/SMZ 200/40 mg per 5mL SUSP
Cefpodoxime 100, 200 mg TAB	Vancomycin 125 mg CAP
Ceftaroline 400, 600 mg INJ	Vancomycin 125mg/2.5mL ORAL SOLN
Ceftazidime 1, 2 g INJ	Vancomycin 1 g INJ
Ceftazidime/avibactam 2.5 g INJ	<b>ANTIFUNGALS</b>
Ceftolozane/tazobactam 1.5 g INJ	Ampho. B conventional 50 mg INJ
Ceftriaxone 1, 2 g INJ	Ampho. B liposomal 50 mg INJ
Cefuroxime 750 mg INJ	Fluconazole 40 mg/mL SUSP
Cephalexin 250, 500 mg CAP	Fluconazole 100 mg TAB
Ciprofloxacin 200, 400 mg INJ	Fluconazole 200, 400 mg INJ
Ciprofloxacin 250, 500 mg TAB	Flucytosine 500 mg CAP
Clarithromycin 500 mg TAB	*Isavuconazonium 186 mg CAP
Clindamycin 150 mg CAP	*Isavuconazonium 372 mg INJ
Clindamycin 600, 900 mg INJ	Itraconazole 100mg CAP
*Colistimethate 150 mg INJ	Micafungin 50, 100 mg INJ
Daptomycin 500 mg INJ	Posaconazole 200 mg/5mL SUSP
Dicloxacillin 250 mg CAP	Posaconazole 300 mg TAB
Doxycycline 100 mg TAB/CAP/INJ	Voriconazole 200 mg INJ/TAB
Ertapenem 1 g INJ	*Voriconazole 40 mg/mL SUSP
Erythromycin 250 mg TAB	<b>ANTIVIRALS</b>
Fidaxomicin 200 mg TAB	Acyclovir 200 mg CAP
*Fosfomycin 3 g SACHET	Acyclovir 200 mg/5ml SUSP
Gentamicin 80mg/2mL INJ - 2mL	Acyclovir 500 mg INJ
Imipenem/cilastatin 250, 500 mg INJ	Acyclovir 800 mg TAB
Levofloxacin 250, 500, 750 mg TAB	Adefovir 10 mg TAB
Levofloxacin 500, 750 mg INJ	Cidofovir 375 mg/5 mL INJ
Linezolid 600 mg TAB/INJ	Entecavir 0.5, 1 mg TAB
Meropenem 500 mg, 1 g INJ	Famciclovir 250 mg TAB
*Meropenem/vaborbactam 1g/1g INJ	Foscarnet 24mg/mL - 500mL INJ
	Ganciclovir 500 mg INJ

§Glecaprevir/pibrentasvir TAB	Lopinavir/ritonavir 200/50 mg TAB
Oseltamivir 30, 75mg CAP	Maraviroc 150 mg TAB
•Peramivir 200 mg INJ	Raltegravir 400, 600 mg TAB
§Velpatasvir/sofosbuvir 100/400 mg TAB	Ritonavir 100 mg TAB
§Velpatasvir/sofosbuvir/voxilaprevir TAB	Tenofovir AF 25 mg TAB
Valacyclovir 500mg TAB	Tenofovir DF 300 mg TAB
Valganciclovir 450mg TAB	Zidovudine 100 mg CAP
<b>ANTIRETROVIRALS</b>	
Abacavir 300 mg TAB	*Zidovudine 50mg/5mL PO syrup
Abacavir 100 mg/5 mL PO SOLN	<b>MISCELLANEOUS ANTI-INFECTIVES</b>
Abacavir/lamivudine 600/300 mg TAB	Albendazole 200mg TAB
Abacavir/lamivudine/dolutegravir 600/300/50 mg TAB	Atovaquone 750mg/5mL SUSP
Abacavir/lamivudine/zidovudine 300/150/300 mg TAB	Atovaquone/proguanil 250/100 mg TAB
Atazanavir 150, 200, 300 mg CAP	Bezlotoxumab 1000 mg/40 mL INJ
•Atazanavir/cobicistat 300/150 mg TAB	*Chloroquine phosphate 500 mg TAB
Bictegravir/emtricitabine/tenofovir AF 50/200/25 mg TAB	Dapsone 100 mg TAB
Darunavir 600, 800 mg TAB	§Ethambutol 100, 400 mg TAB
•Darunavir 100 mg/mL PO SUSP	§Hydroxychloroquine 200 mg TAB
Darunavir/cobicistat 800/150 mg TAB	Isoniazid 100, 300 mg TAB
Dolutegravir 50 mg TAB	Ivermectin 3mg TAB
Dolutegravir/lamivudine 50/300 mg TAB	Miconazole 2% topical POWDER
Dolutegravir/rilpivirine 50/25 mg TAB	*Nitazoxanide 500mg TAB
Doravirine 100 mg TAB	*Nystatin 100,000 units/g topical POWDER
Efavirenz 200 mg CAP	Pentamidine 300mg INJ/INHALATION
Efavirenz 600 mg TAB	*Praziquantel 600 mg TAB
Efavirenz/lamivudine/tenofovir DF 600/300/300 TAB	Primaquine 26.3 mg TAB
Elvitegravir/cobicistat/emtricitabine/tenofovir AF 150/150/200/10 mg TAB	Pyrazinamide 500 mg TAB
Elvitegravir/cobicistat/emtricitabine/tenofovir DF 150/150/200/300 mg TAB	Rifabutin 150 mg CAP
Emtricitabine 200 mg CAP	Rifampin 300 mg CAP
Emtricitabine/tenofovir AF 200/25 mg TAB	Rifampin 600 mg INJ
Emtricitabine/tenofovir DF 200/300 mg TAB	Rifapentine 150 mg TAB
Emtricitabine/rilpivirine/tenofovir AF 200/25/25 mg TAB	*Rifaximin 200 mg TAB
Emtricitabine/rilpivirine/tenofovir DF 200/25/300 mg TAB	Terbinafine 250 mg TAB
Enfuvirtide 90 mg INJ	*Tinidazole 500 mg TAB
Etravirine 200 mg TAB	Tolnaftate 1% topical POWDER
Lamivudine 100, 150 mg TAB	
Lamivudine 50 mg/5 mL PO SOLN	
Lamivudine/zidovudine 150/300 mg TAB	

Shaded background = Restricted to ID

• = Nonformulary; requires "Prior authorization drug request" consult

§ = Restricted to ID/GI and requires PADR consult

## **IMPORTANT PHONE EXTENSIONS**

Inpatient Pharmacy	x6420
Outpatient Pharmacy	x6767
General Chemistry	x6498
Microbiology Lab	x6501
Infection Prevention/Infection Control	x4827

For questions/comments, please contact ID PharmD:

Jaela Fredenrich	x4236
Amanda Mercurio	x6035
PGY2 Infectious Diseases Pharmacy Resident	x5822

For Infectious Diseases Consult or  
Antimicrobial Stewardship, please refer to the  
Infectious Diseases call schedule on the Sharepoint



James A. Haley Veterans' Hospital  
13000 Bruce B. Downs Blvd  
Tampa, FL 33612  
813-972-2000

**Created/Maintained by:**

Amanda K. Mercurio, PharmD, BCPS  
Clinical Pharmacy Specialist - Infectious Diseases